

Fig. 1

Compound A

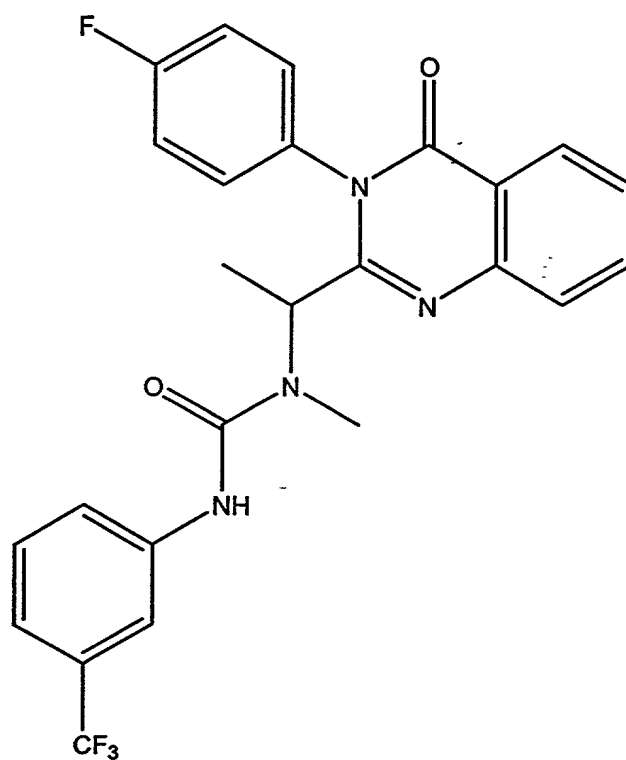


Fig. 2A

Compound B

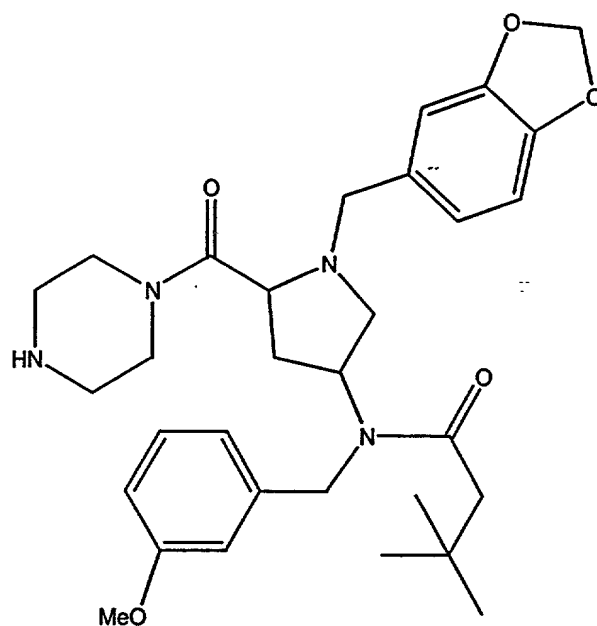
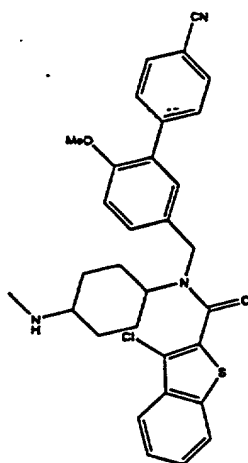


Fig. 2B



Agonist Z

Fig. 3

Gli-1, Gene Expression in the Lung

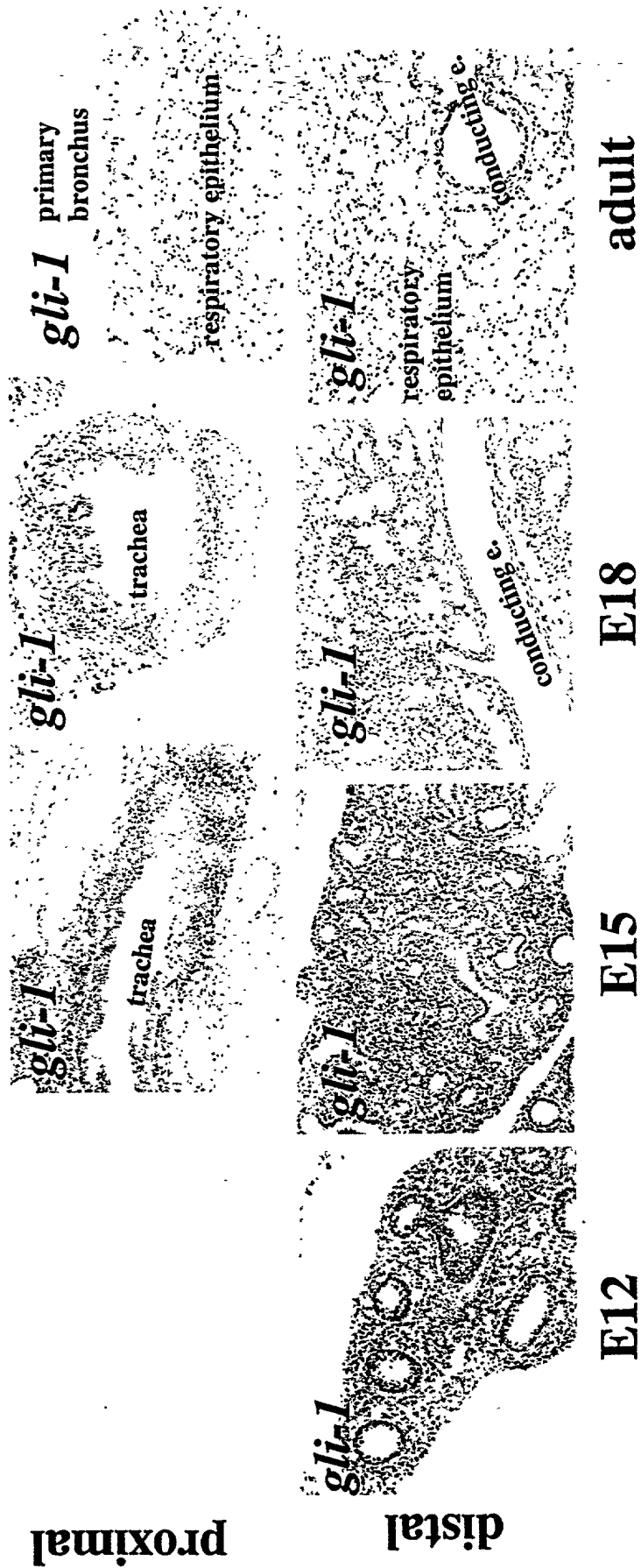


Fig. 4

Gli-1 expression is inversely related to lung maturation

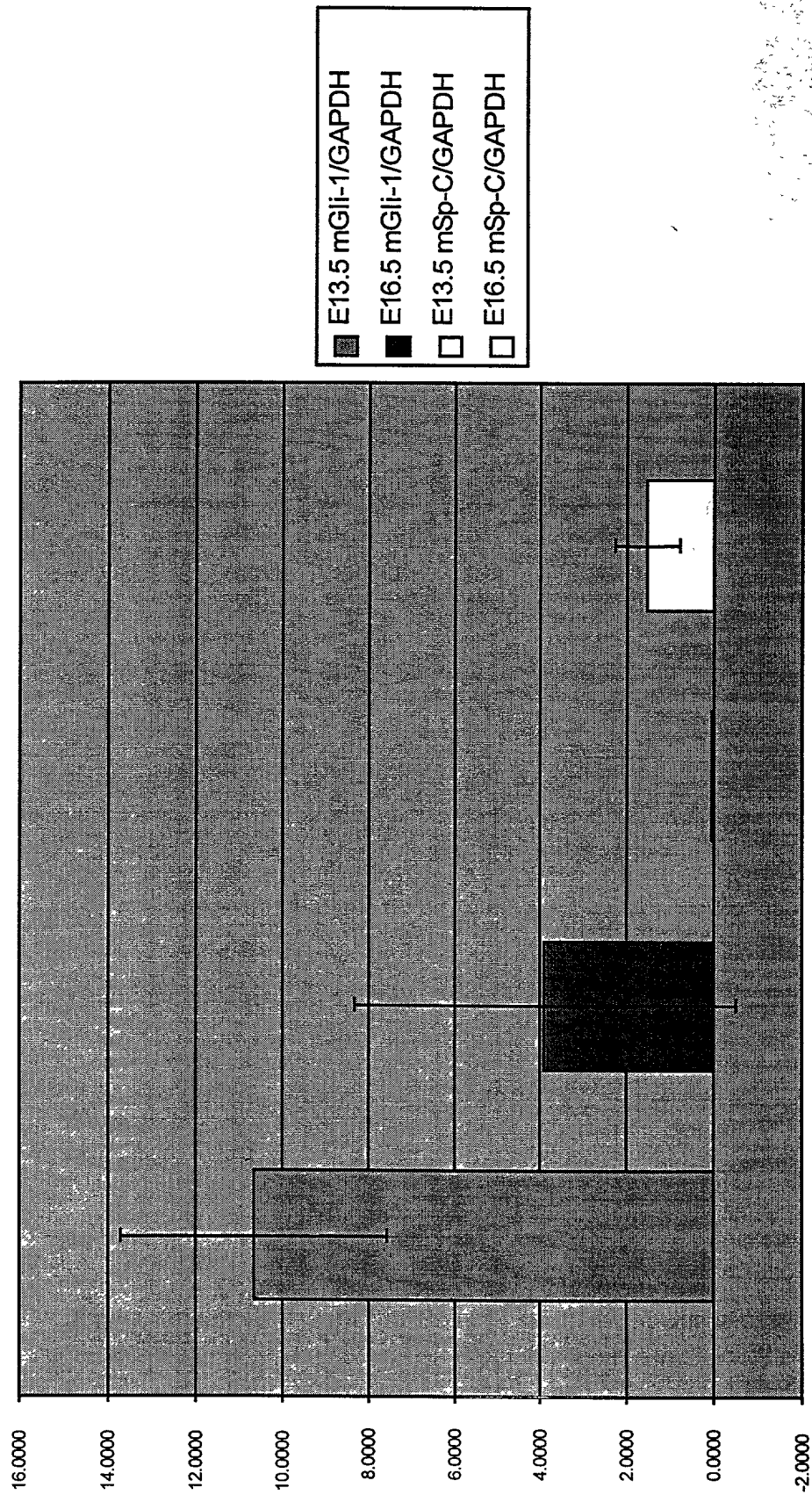


Fig. 5

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Compound B Downregulates mGli-1 in Lung Explant Cultures

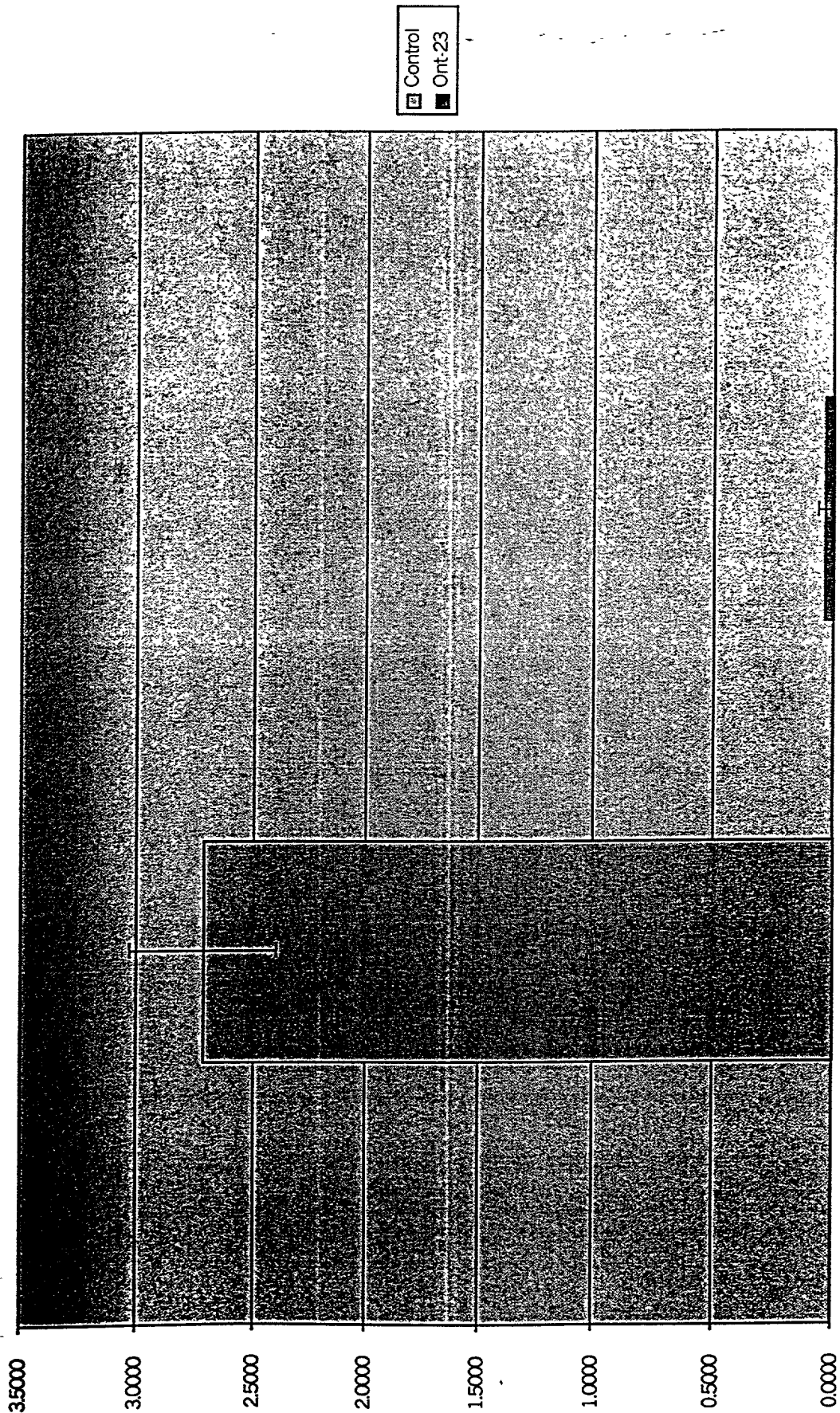


Fig. 6

Compound B Treatment Increases Surfactant type C Production in Embryonic Mouse Lungs

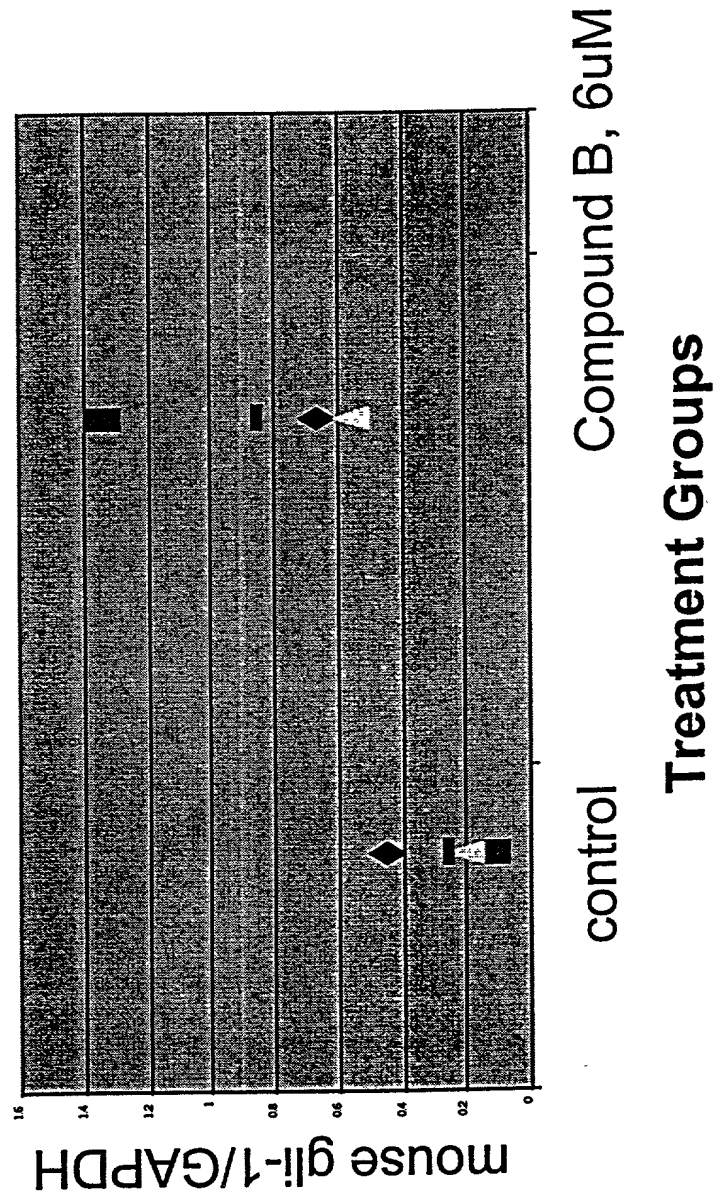
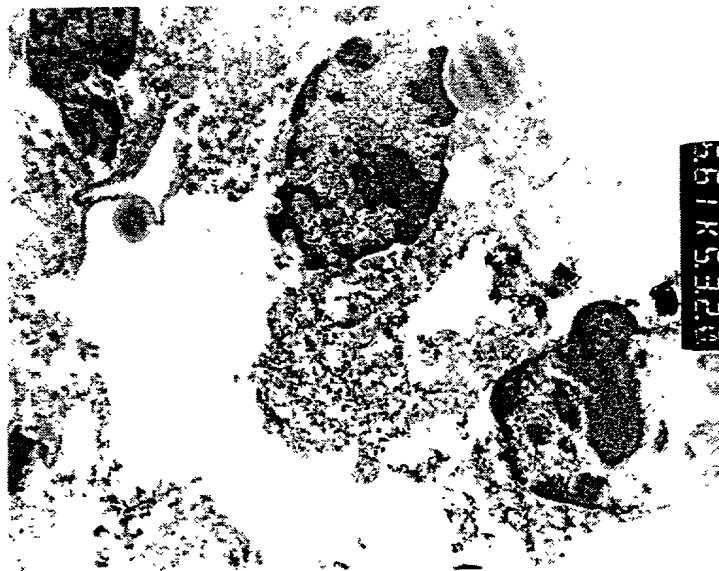


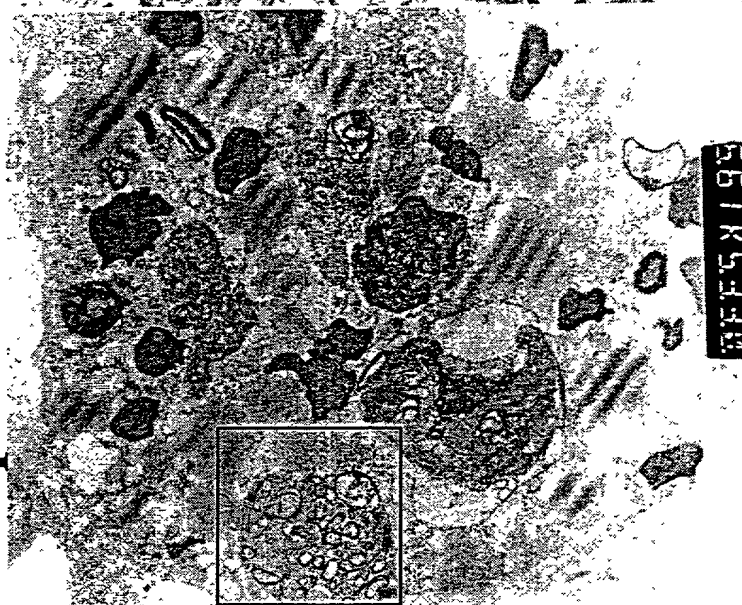
Fig. 7

Type II Pneumocytes in Compound B Treated Lung Cultures Differentiate Prematurely

vehicle



Compound B



Compound B, inset

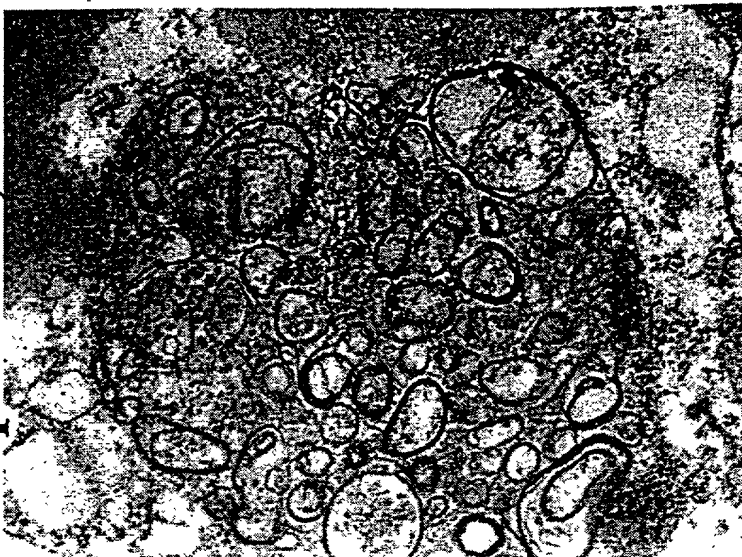


Fig. 8

Compound B

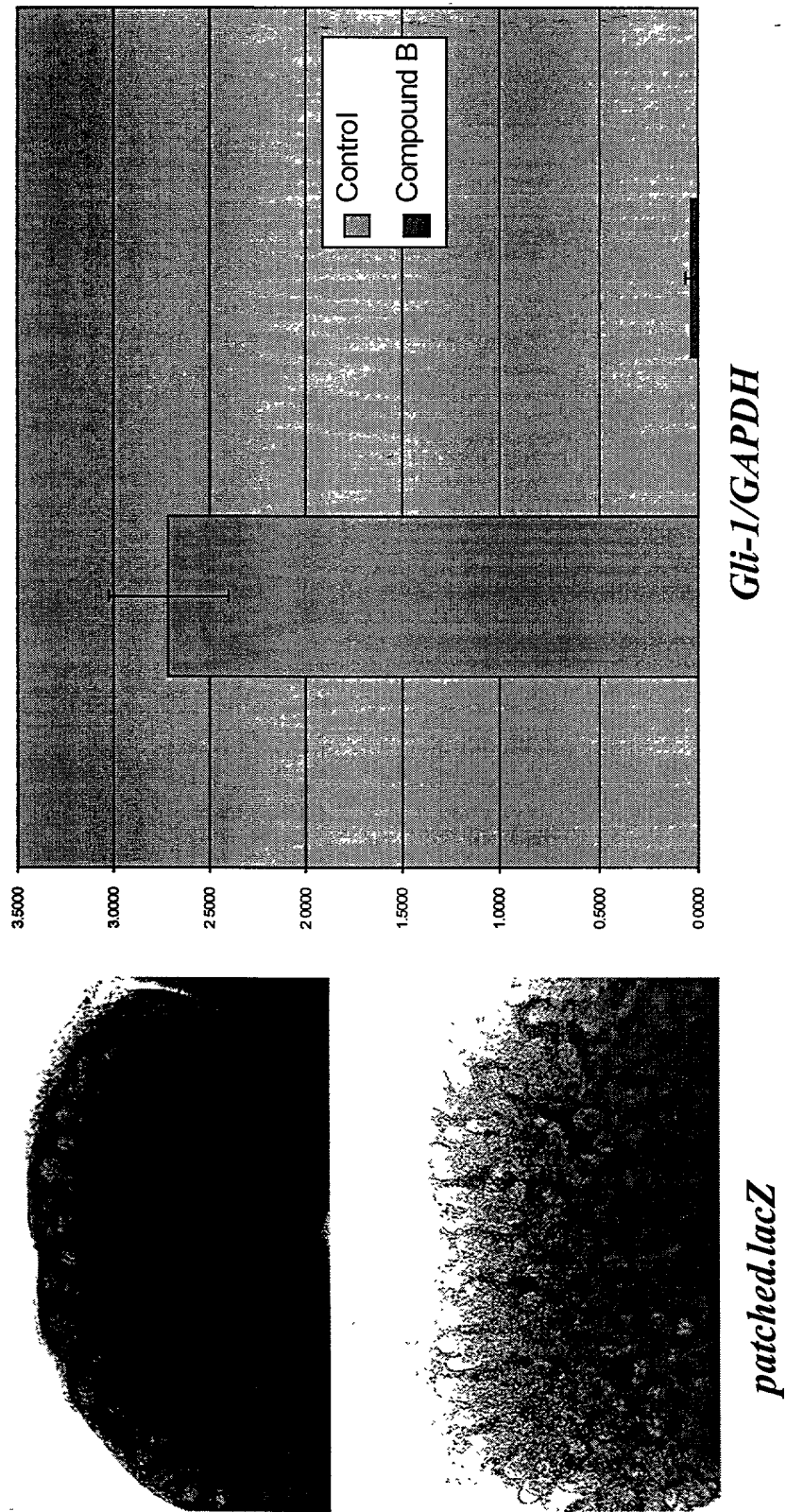


Fig. 9

Compound B

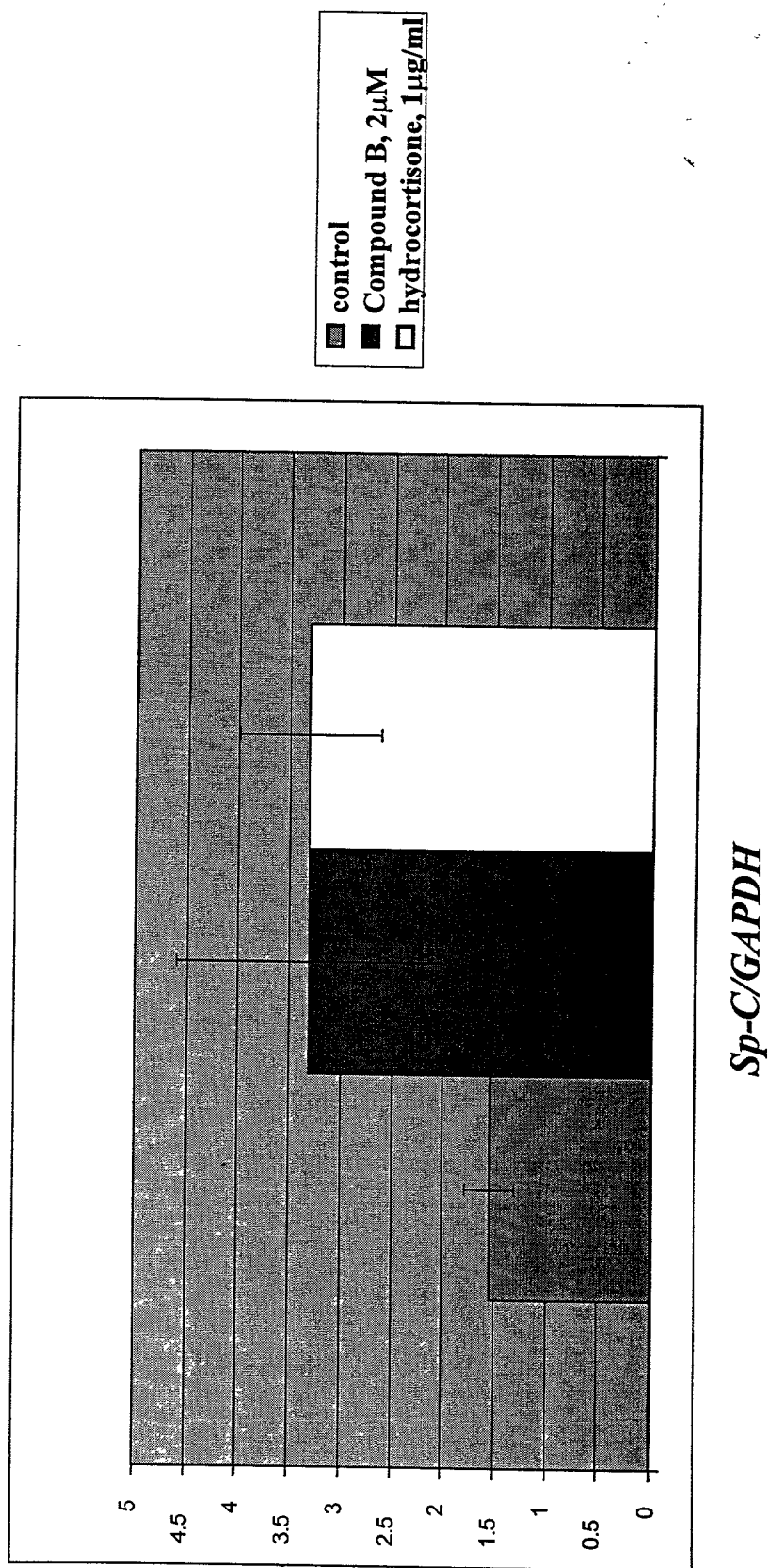


Fig. 10

Shh Protein and Hedgehog Agonist Z

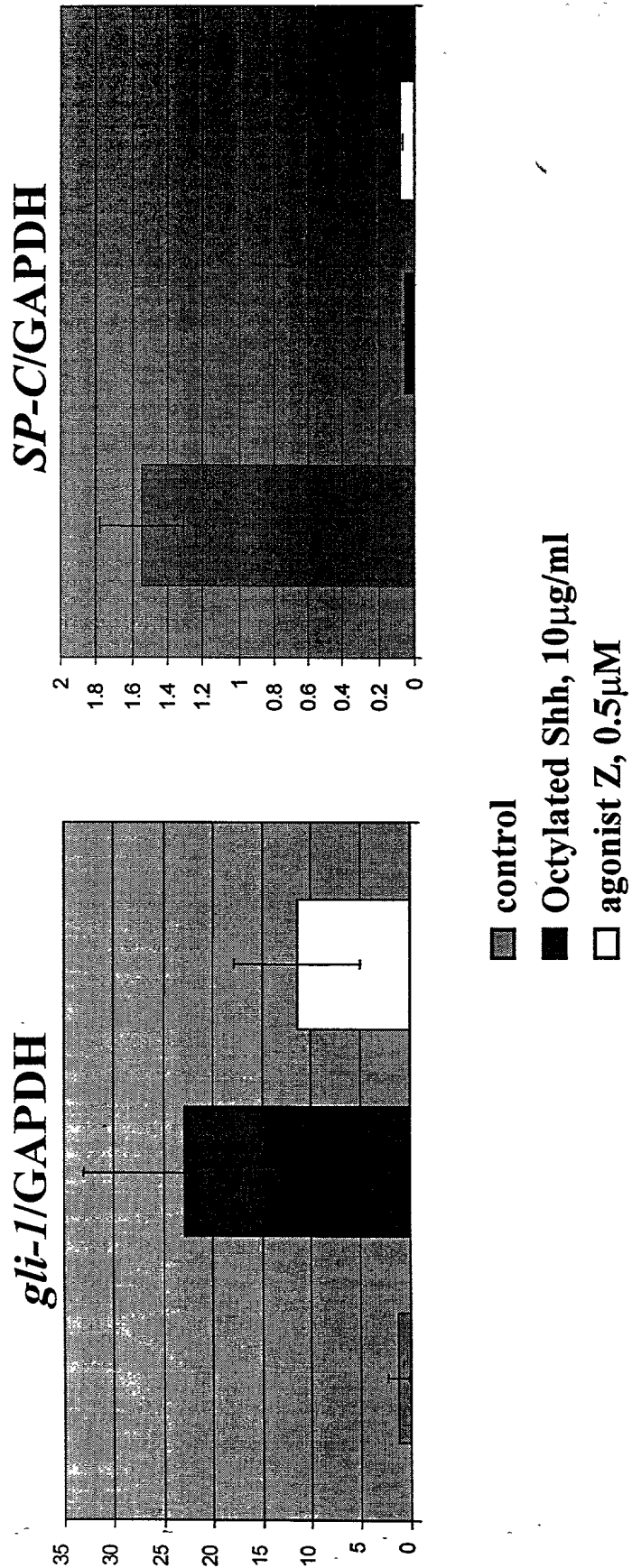


Fig. 11

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Breast Cancer *Gli-1* In Situ Hybridization - Epithelial Expression

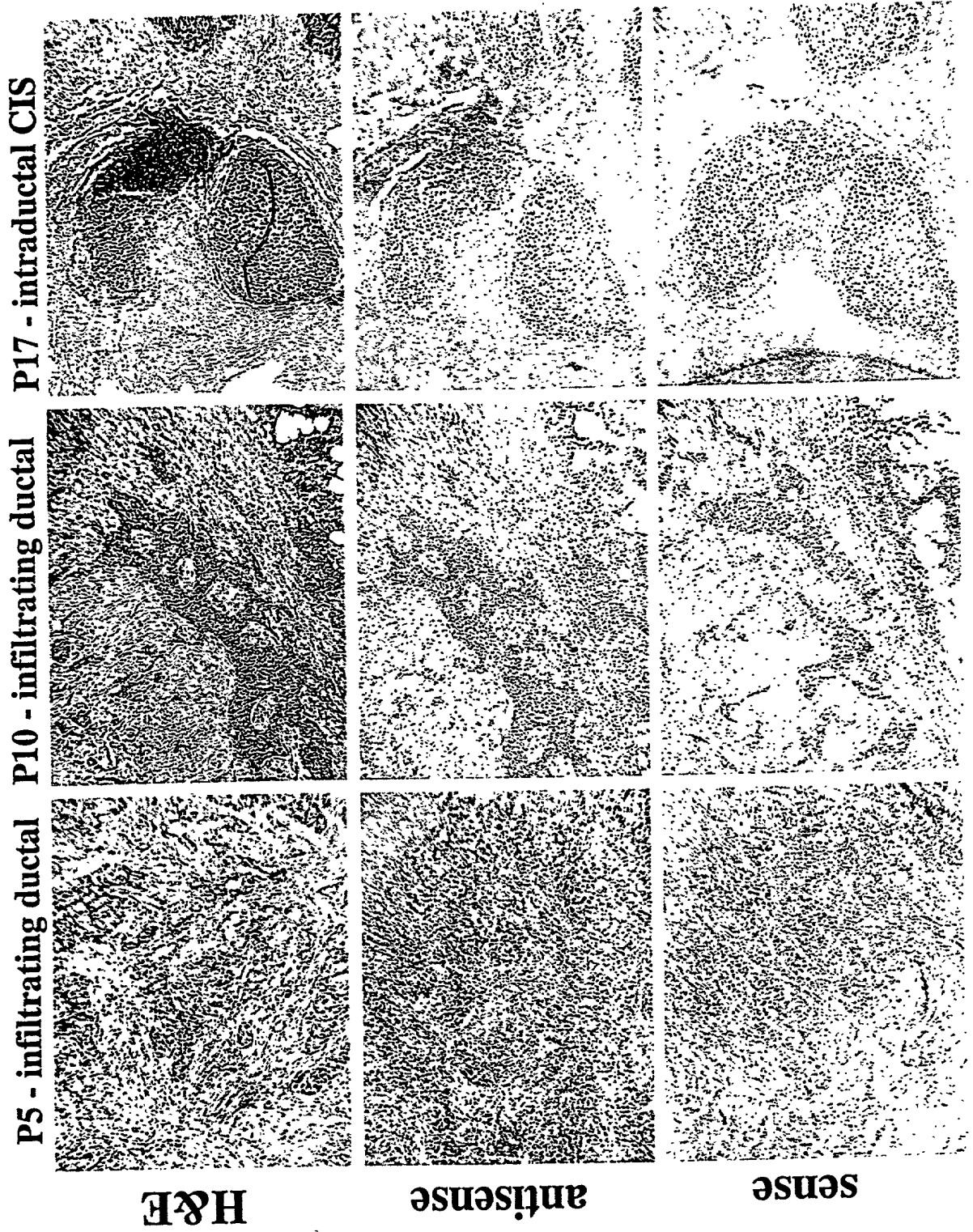


Fig. 12

Lung Cancer *Gli-1* In Situ Hybridization

L4 - adenocarcinoma L17 - small cell carcinoma L20 - small cell carcinoma

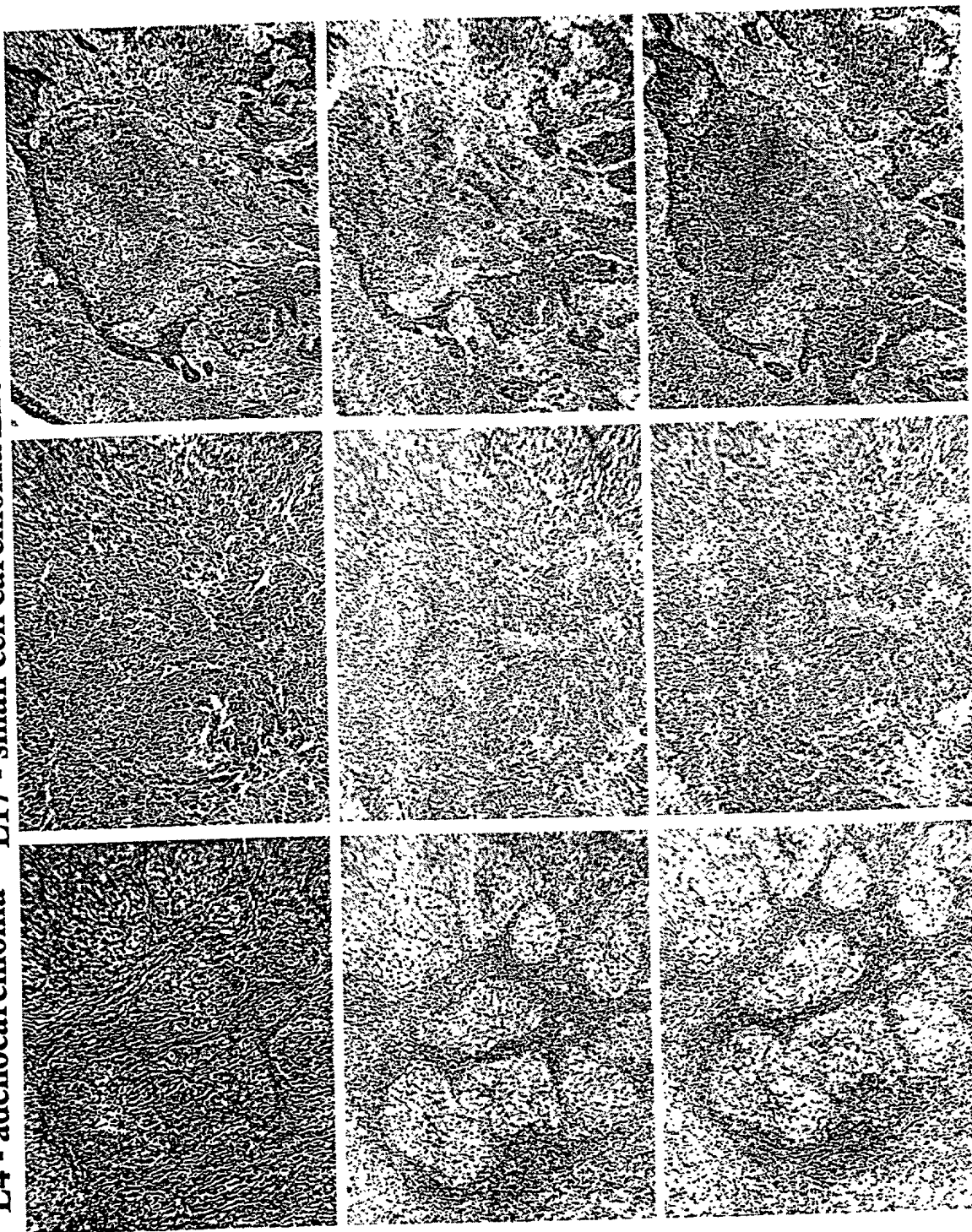
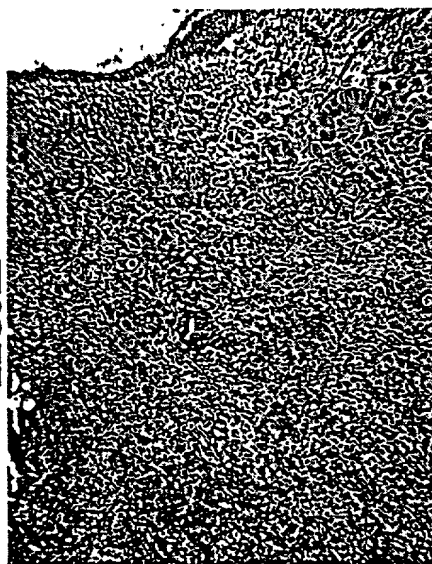


Fig. 13

Prostate Cancer *Gli-1* In Situ Hybridization - Stromal Expression

H&E



antisense



sense



Fig. 14

FOOT 492266

BPH *Gli-1* In Situ Hybridization - Stromal Expression

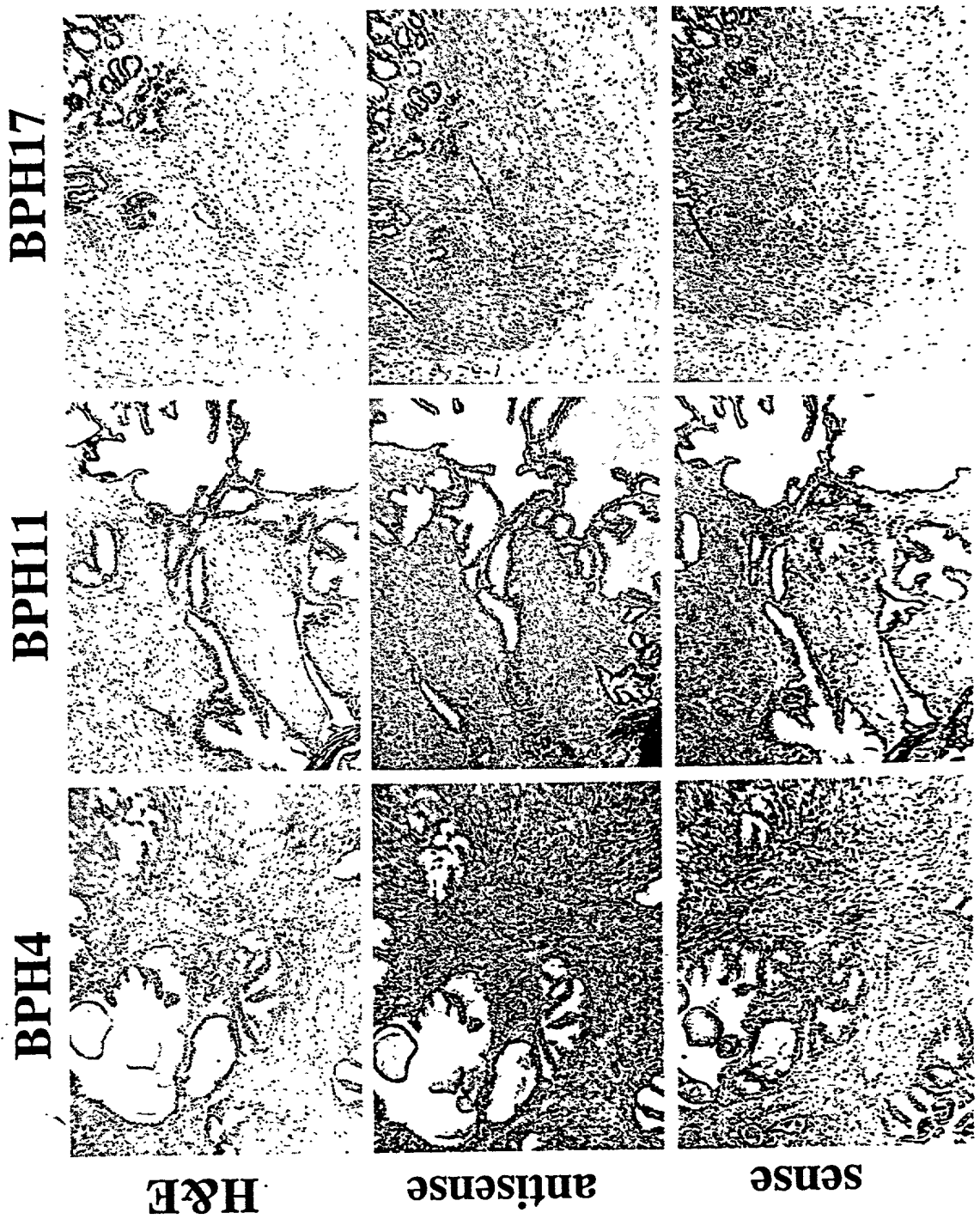


Fig. 15

Hedgehog Signaling in Mouse Bladder

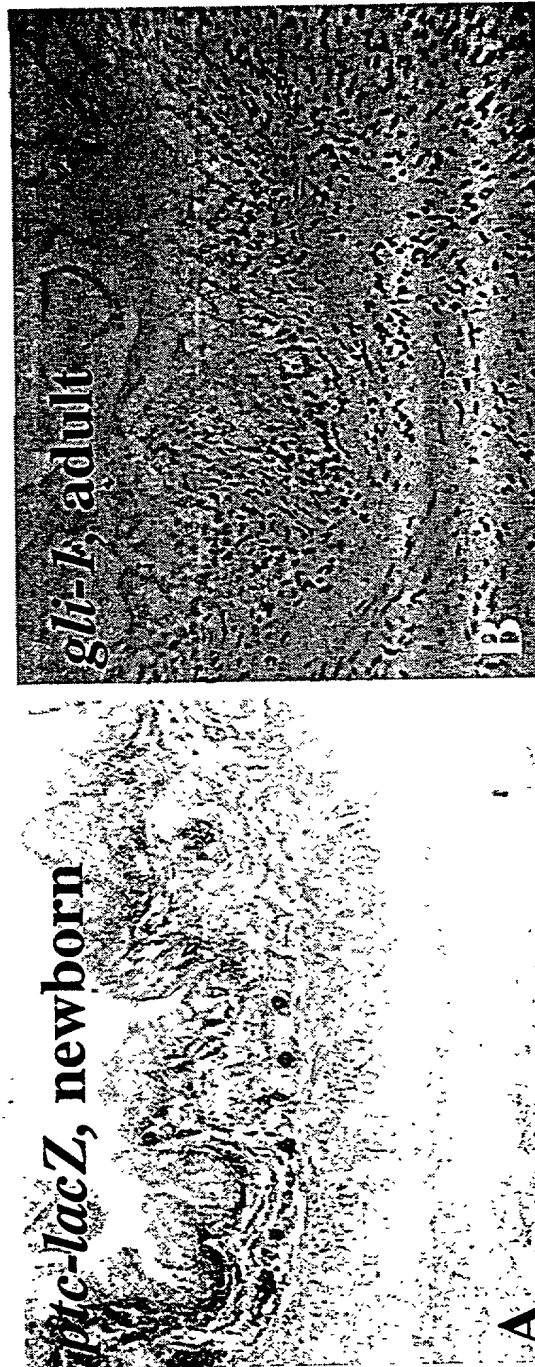


Fig. 16

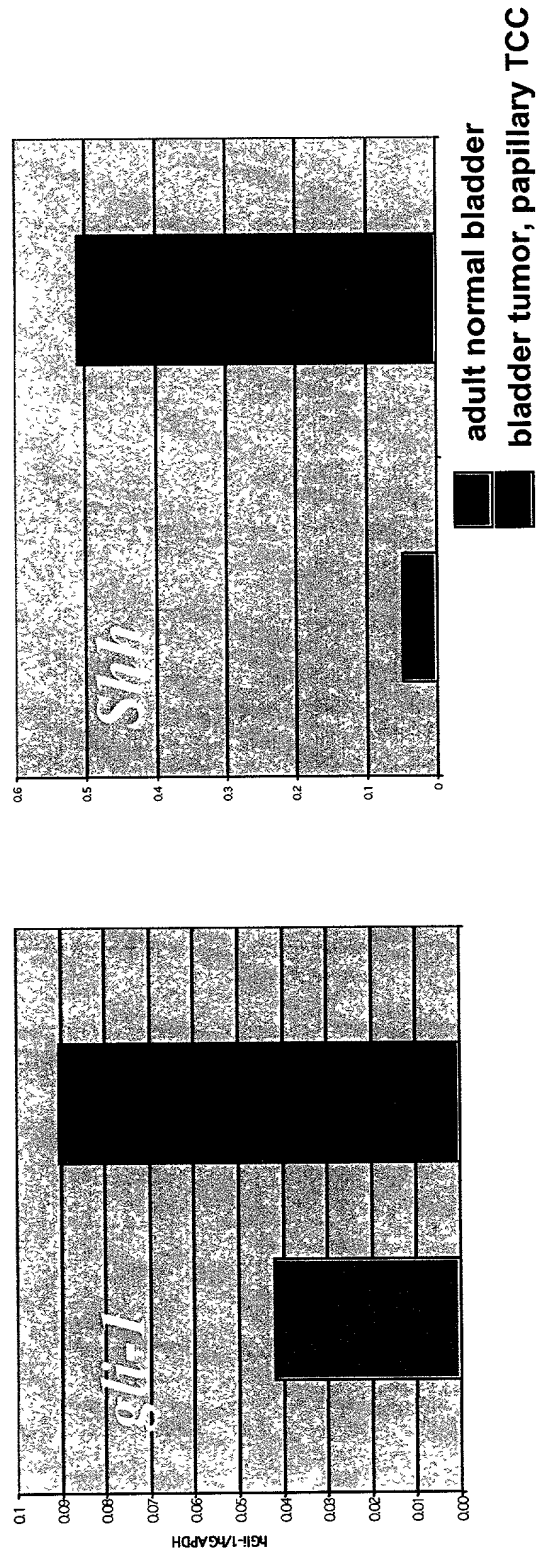
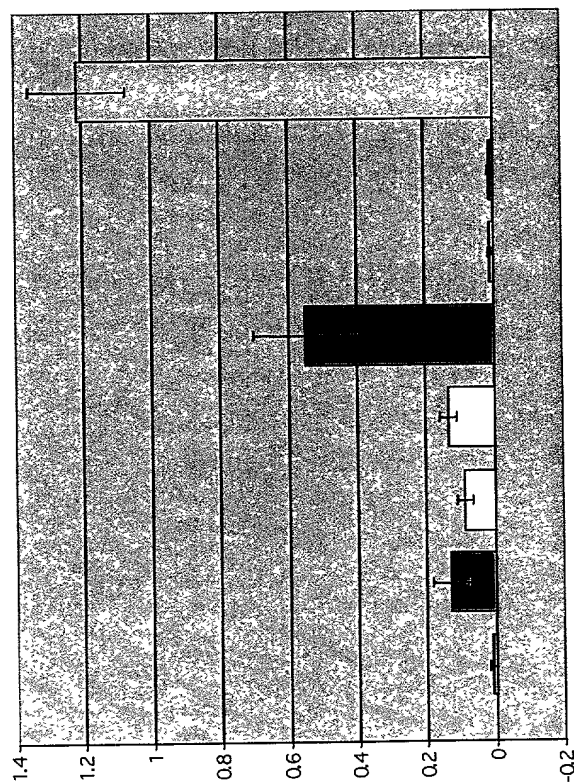


Fig. 17

HH SIGNALING IN BLADDER CANCER CELL LINES

(1d in 10% FBS, 2d in 1% FBS)

hShh



hGli-1

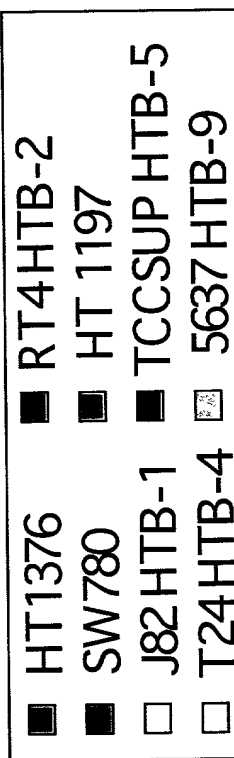
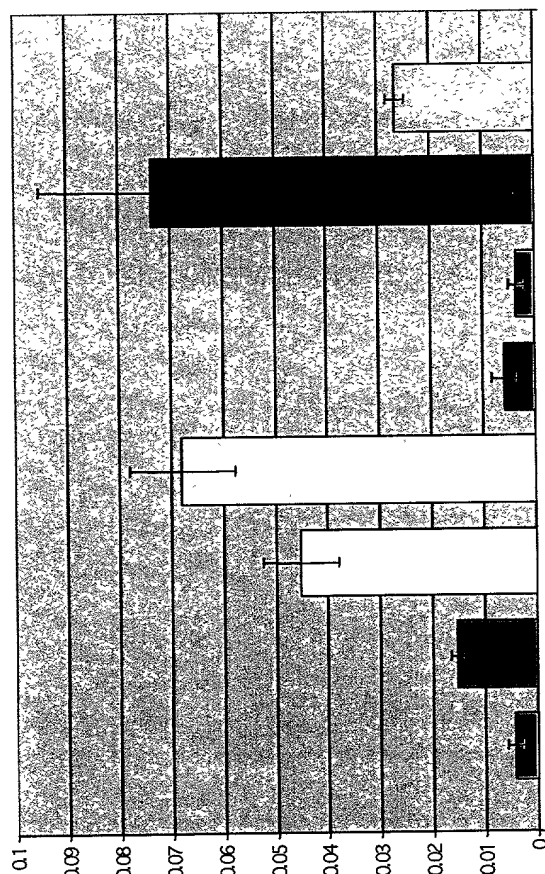


Fig. 18

HH SIGNALING IN BLADDER CANCER CELL LINES

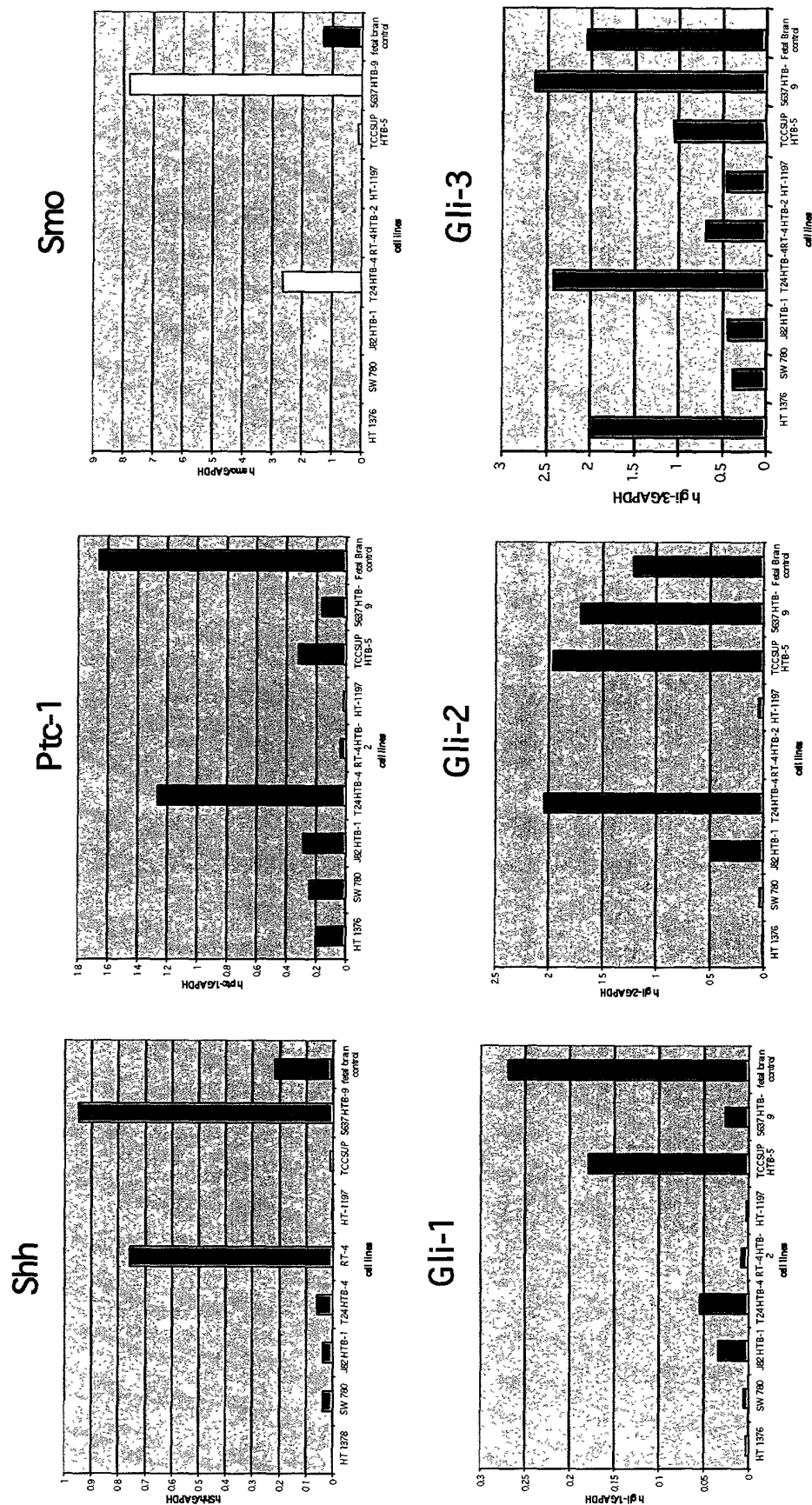


Fig. 19

IN VITRO EFFICACY

Gli-luc Assay

A. S12 fibroblast cell line with luciferase reporter

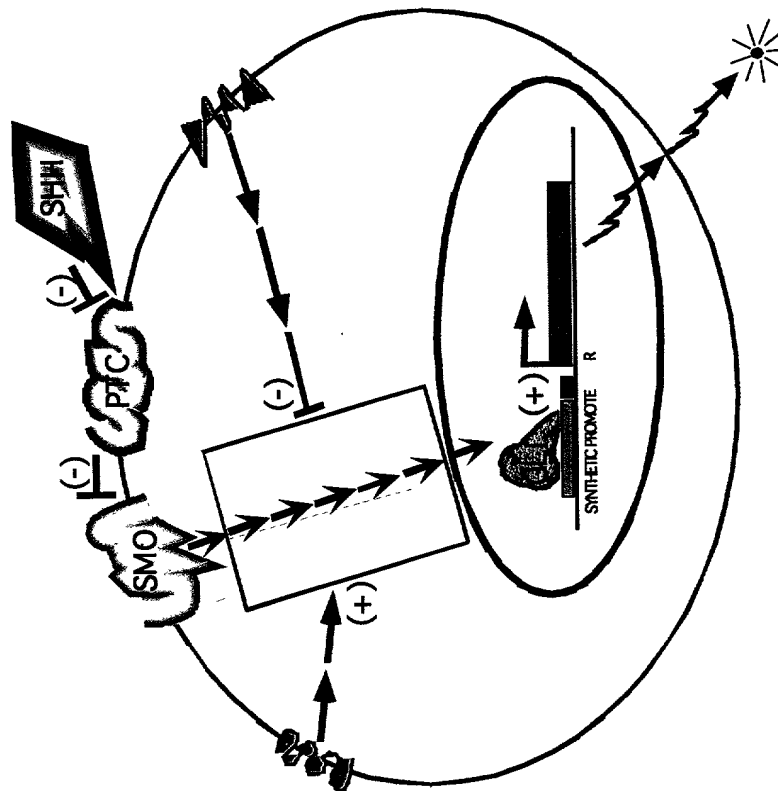
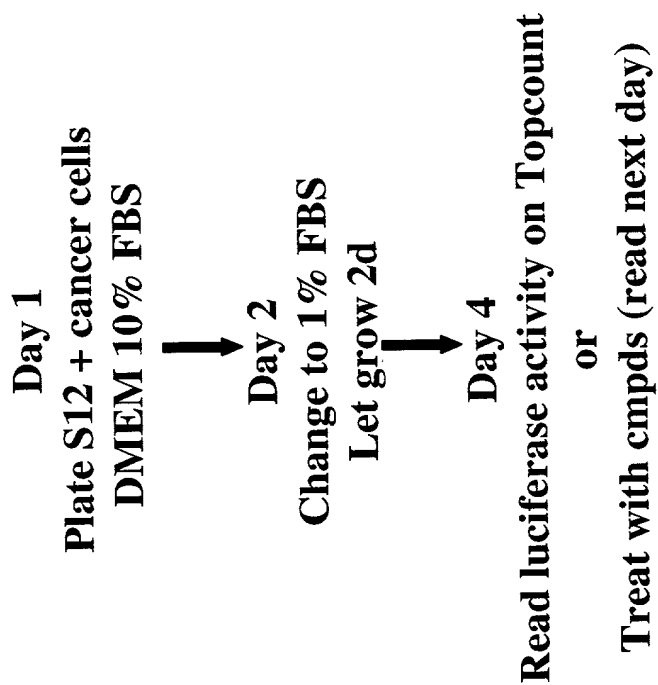


Fig. 20

B. Flow-chart



GLI-LUC ASSAY ON BLADDER CANCER CELL LINES

(S12 + cancer cell co-cultures, 1d in 10% FBS, 2d in 1% FBS)

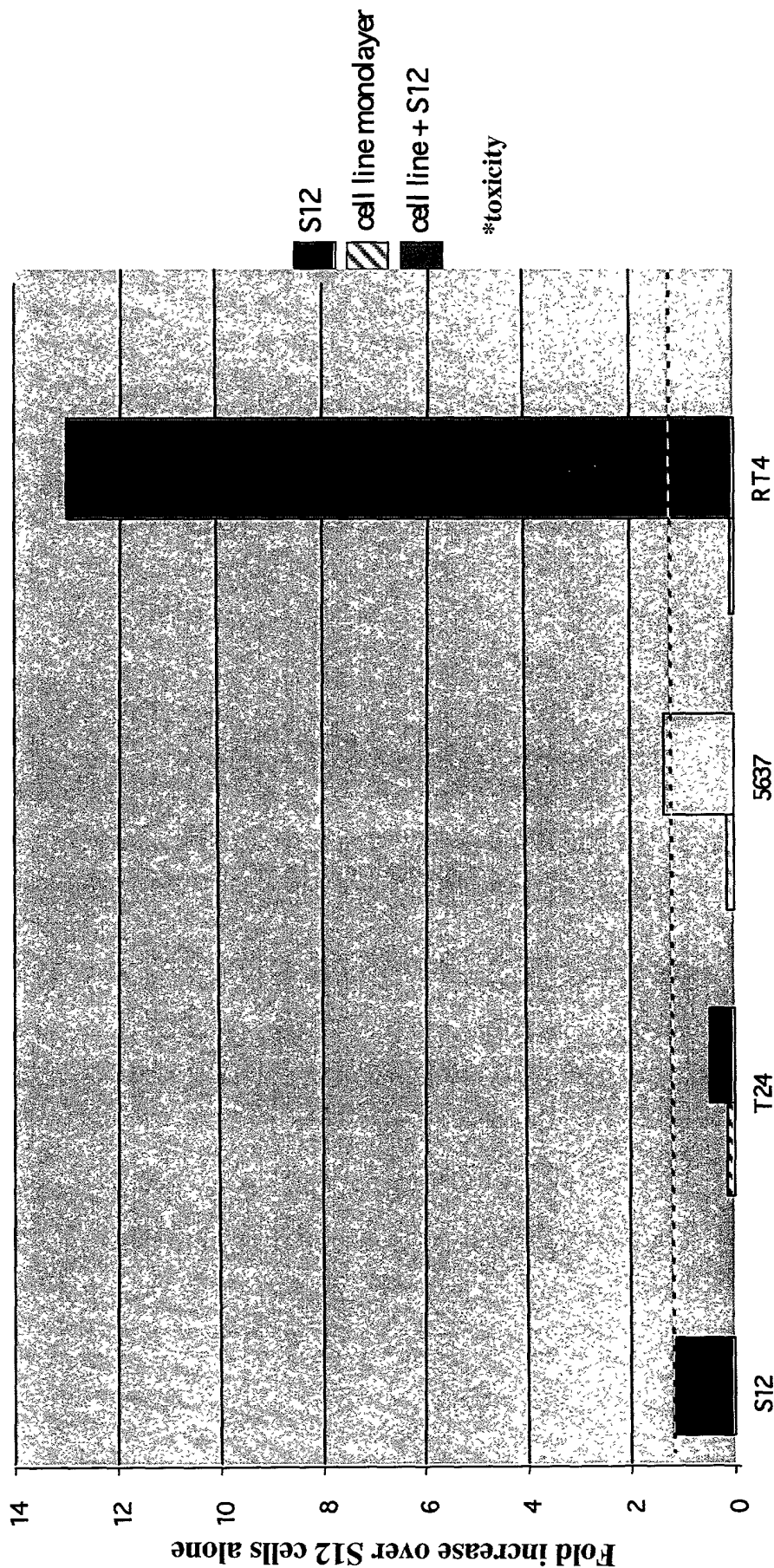
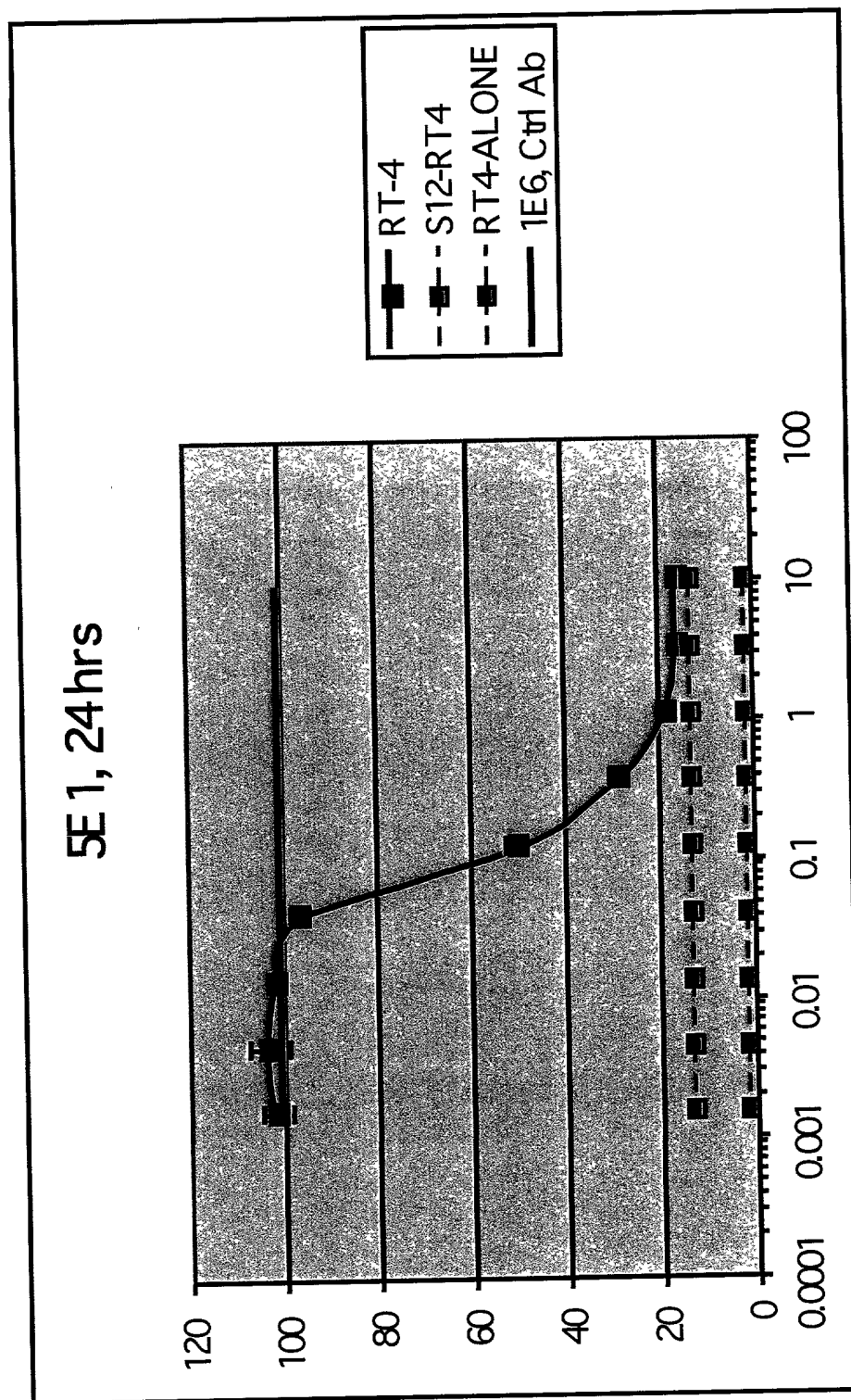


Fig. 21

GLI-LUC ASSAY ON RT-4



IC₅₀=85ng/ml IC₉₀=500ng/ml

Fig. 22

RT-4 IN VIVO EFFICACY EXPERIMENT

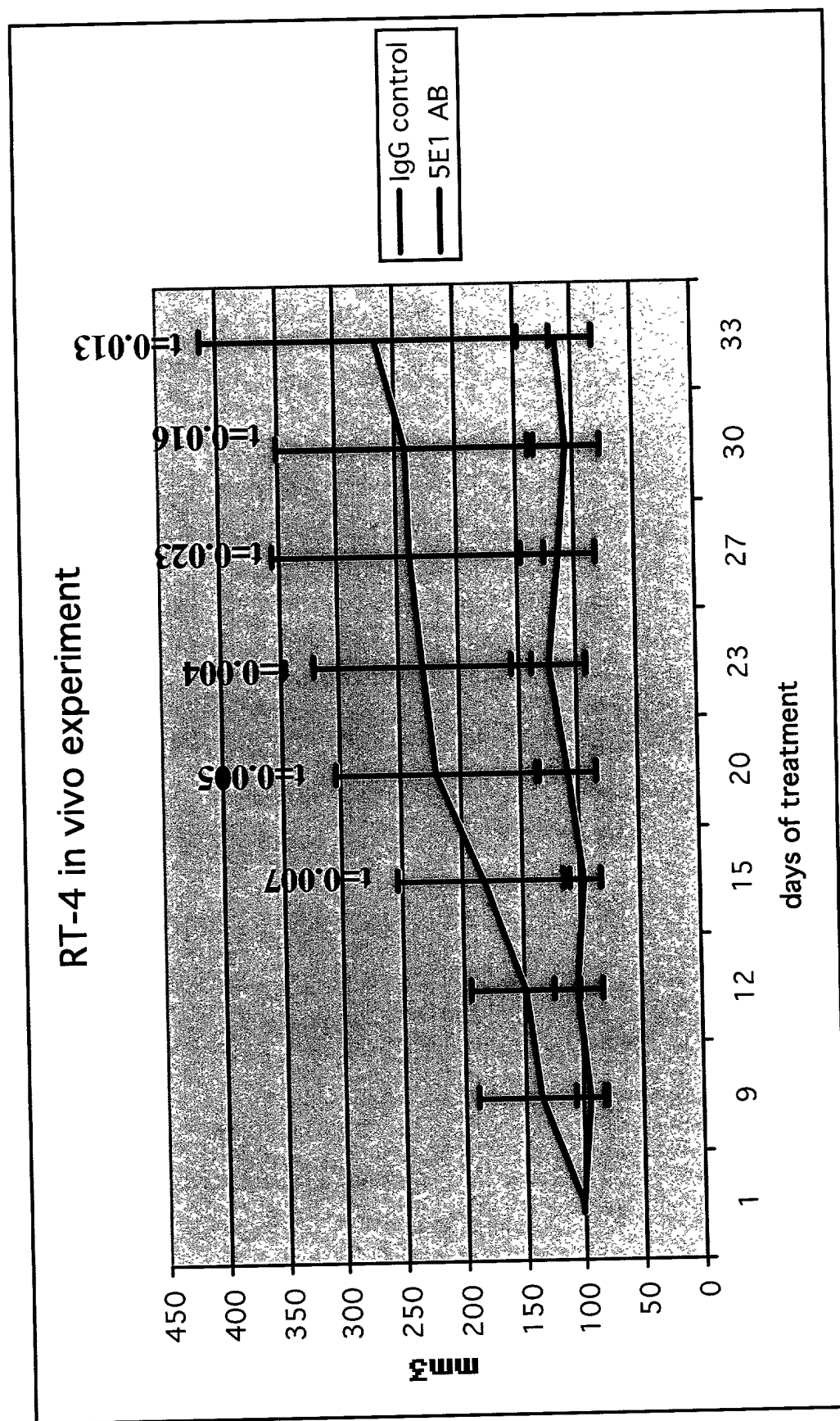
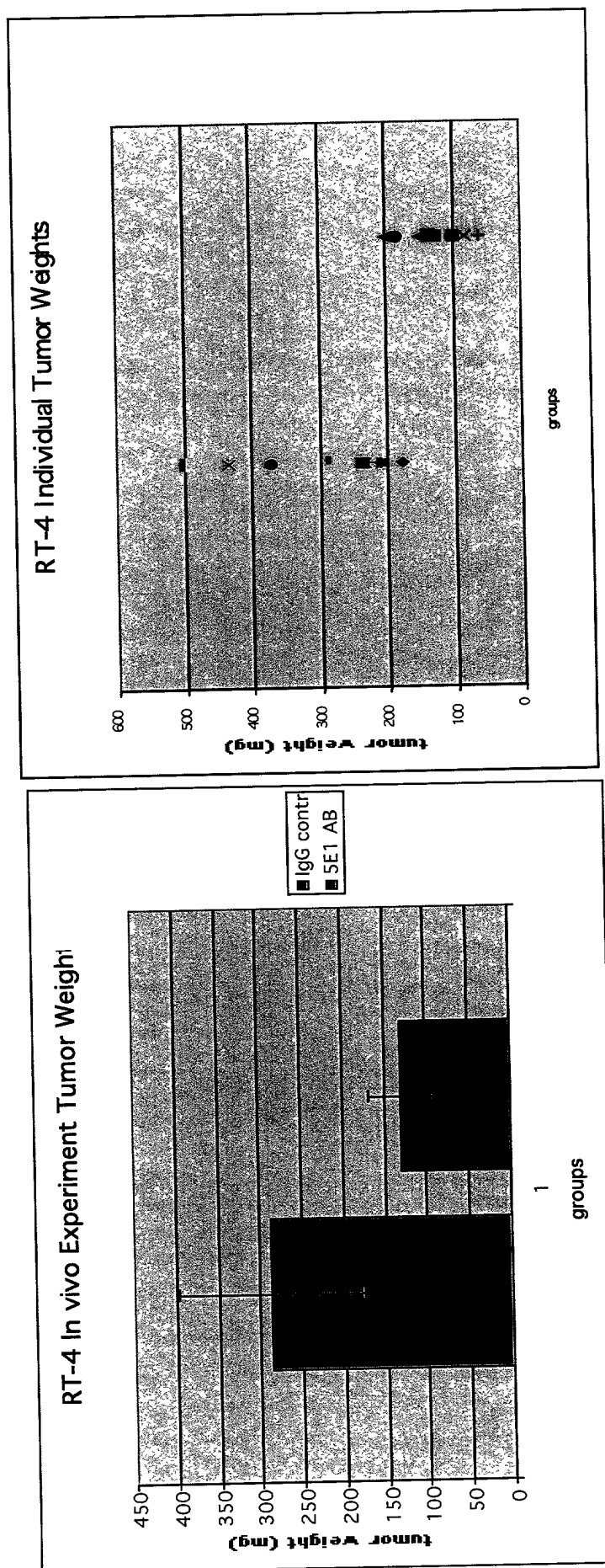
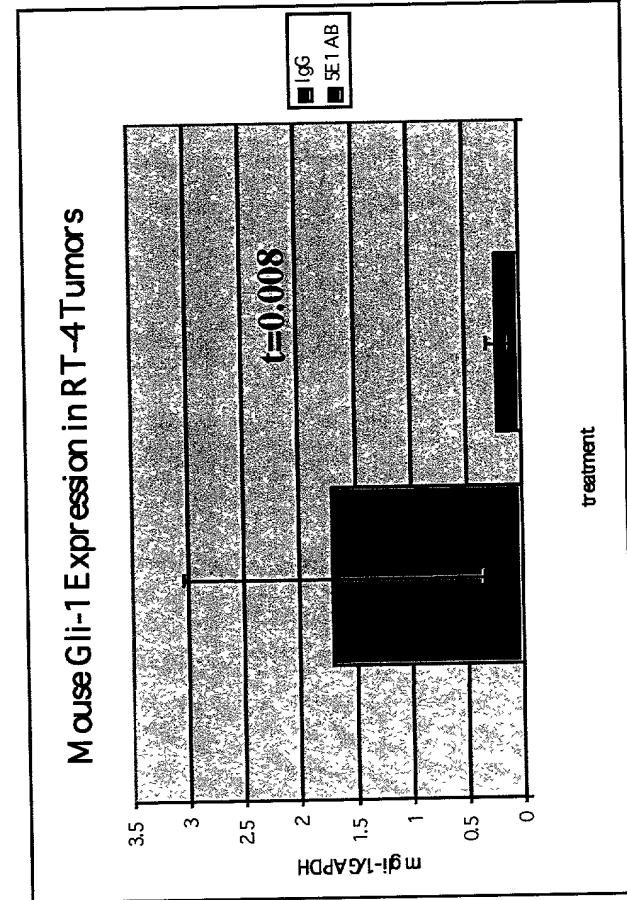


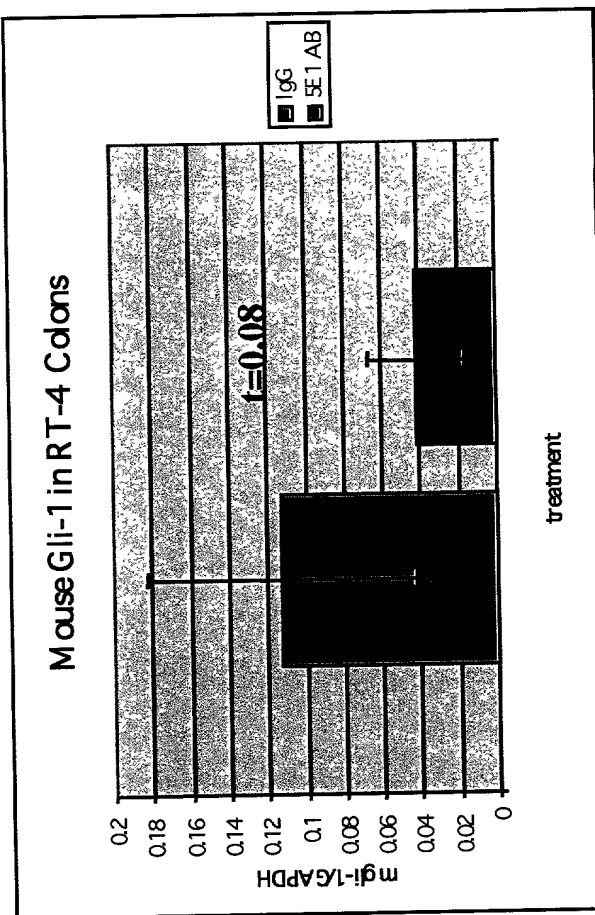
Fig. 23



Mouse Gli-1 Expression in 5E1-Treated RT-4 Tumors



mouse small intestine standard = 0.06



mouse small intestine standard = 0.1

Fig. 25

PROSTATE CANCER

Shh is Expressed in Prostate Cancer

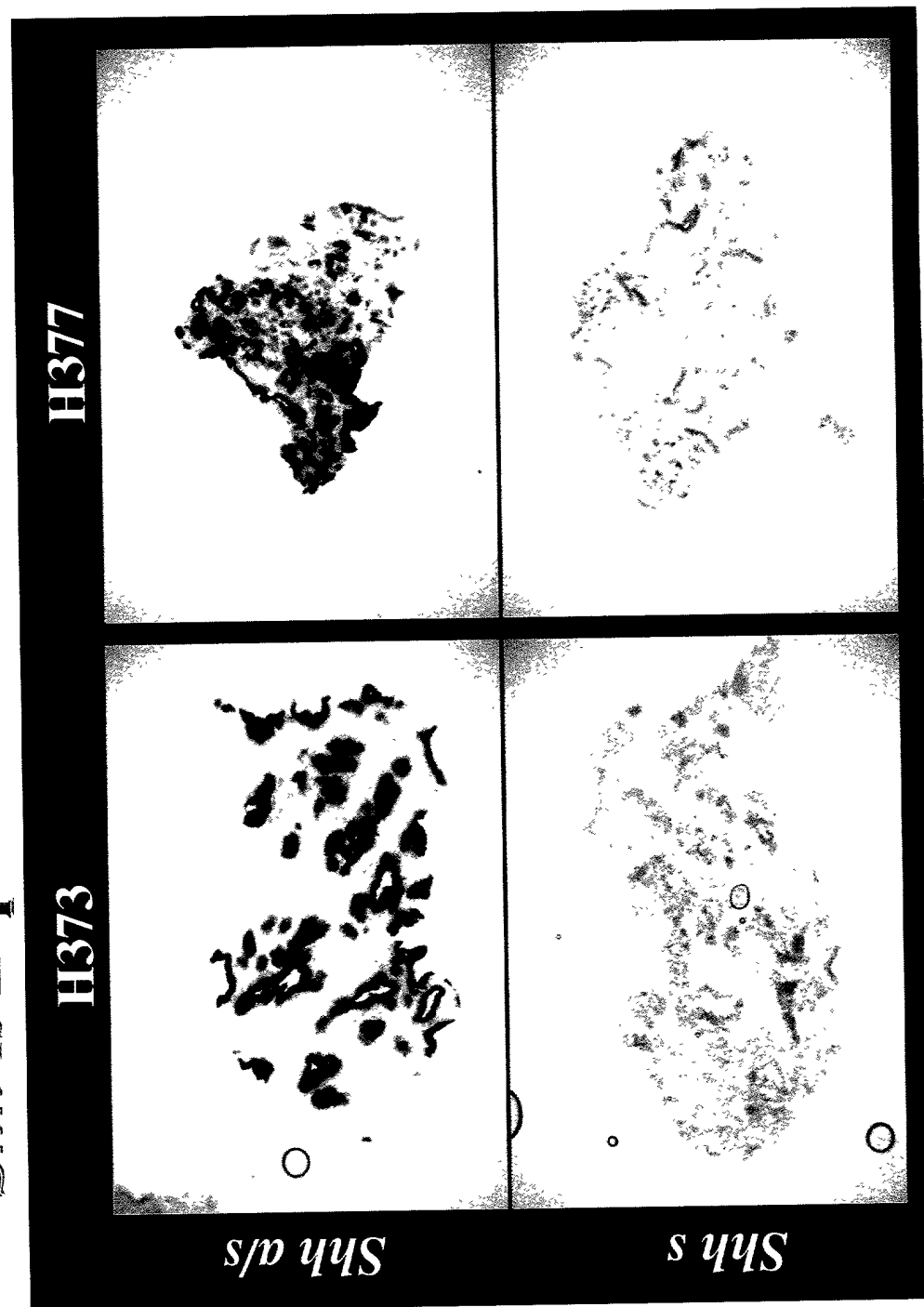


Fig. 26

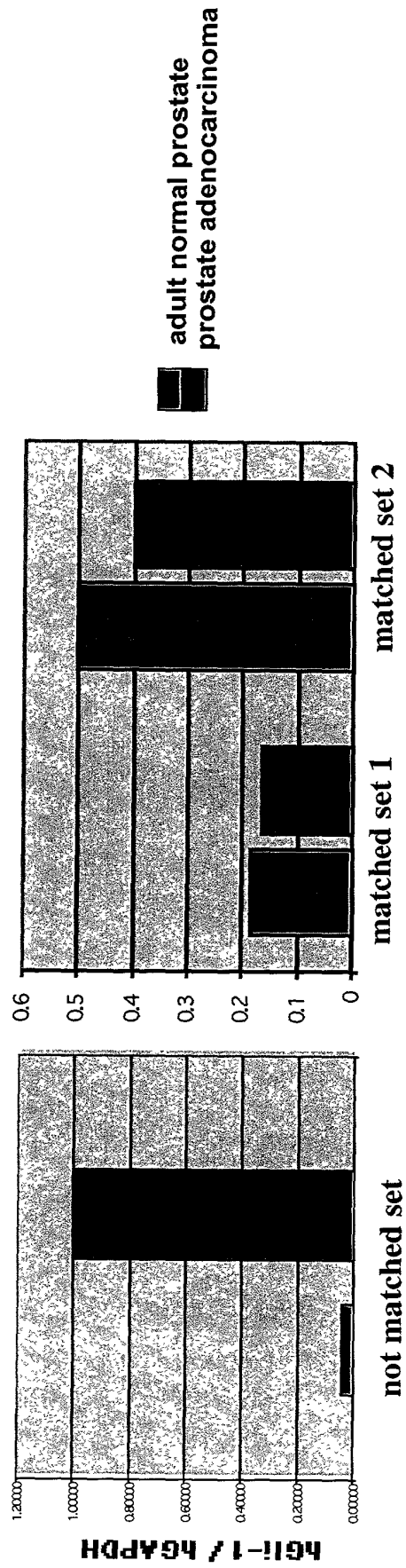


Fig. 27

HH SIGNALING IN PROSTATE CANCER CELL LINES

(1d in 10% FBS, 2d in 1% FBS)

hGli-1

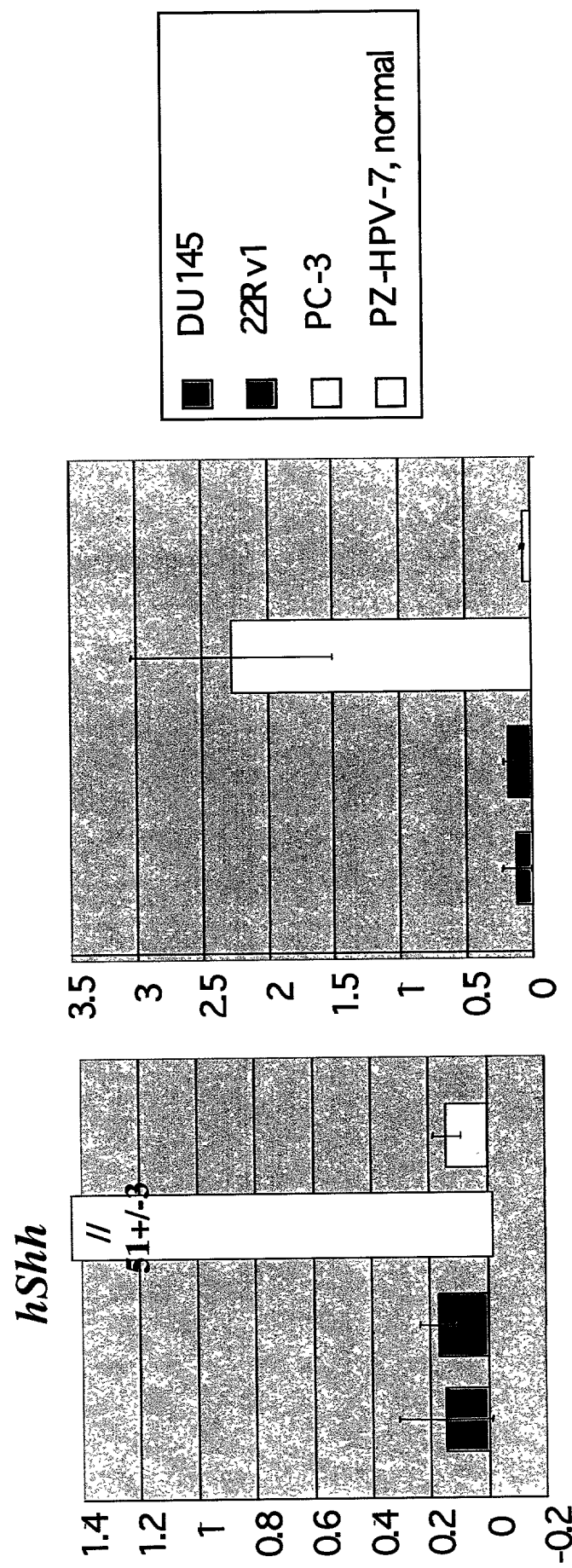


Fig. 28

GLI-LUC ASSAY ON PROSTATE CANCER CELL LINES

(S12 + cancer cell co-cultures, 1d in 10% FBS, 2d in 1% FBS)

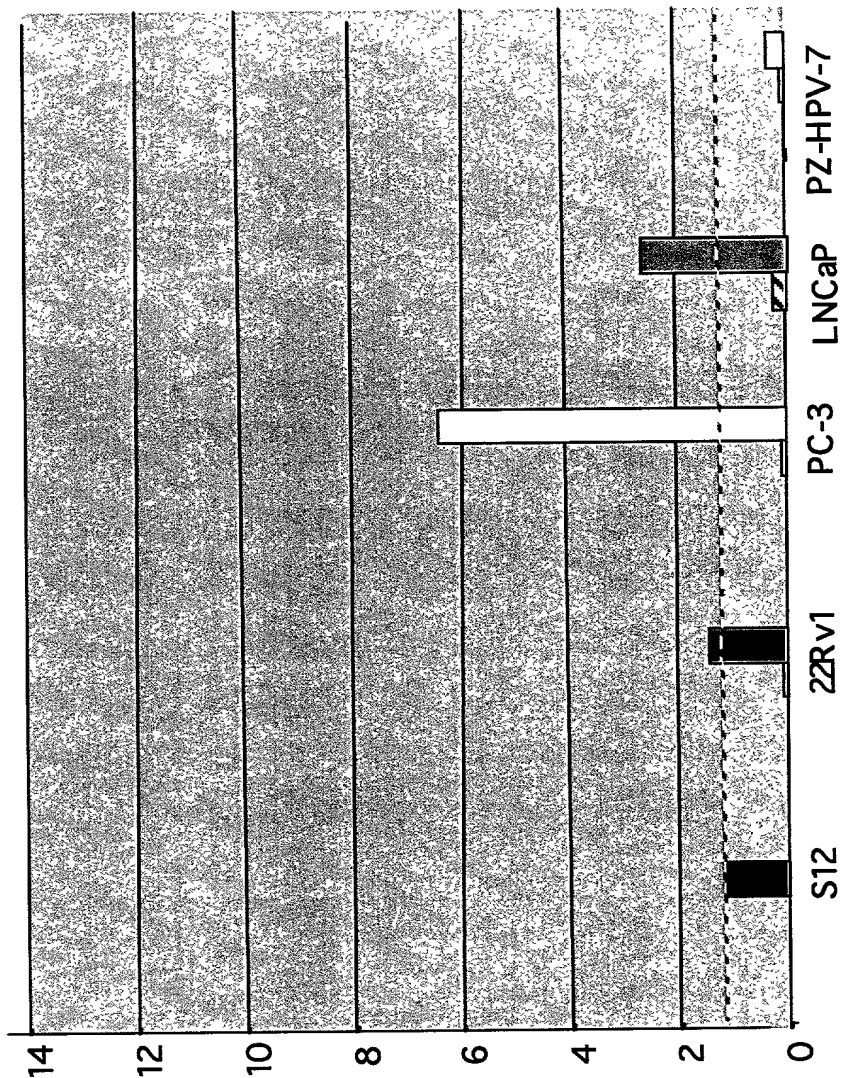


Fig. 29

IN VITRO EFFICACY

Inhibition of hedgehog signaling by Hedgehog Antagonists (Gli-luc, 24 hrs)

5E1

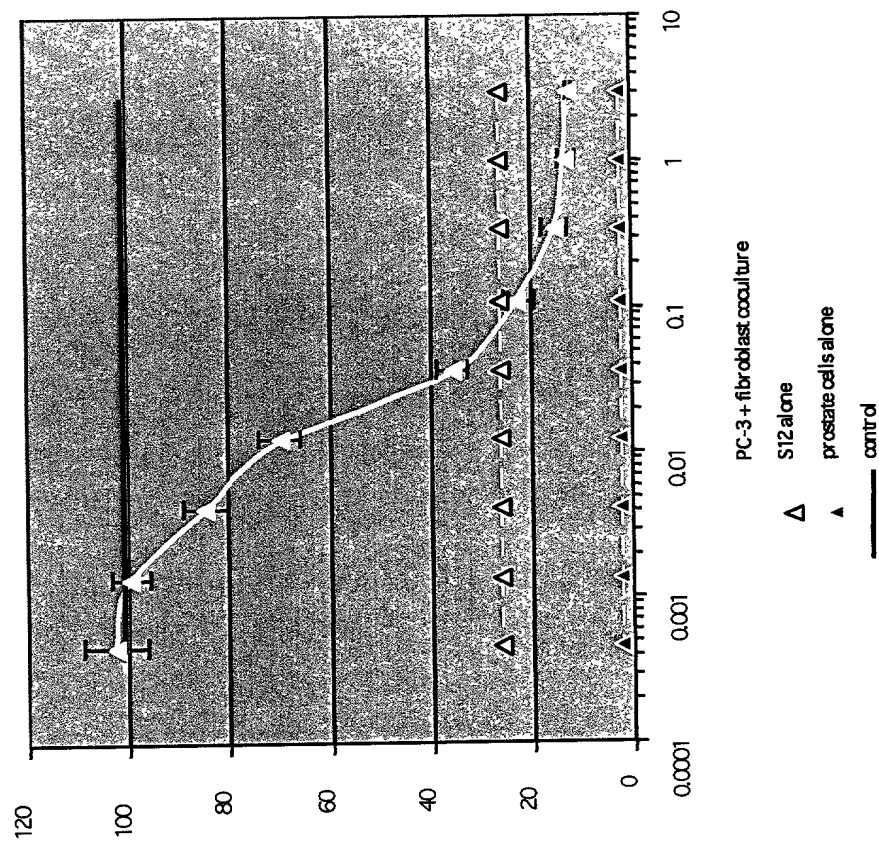
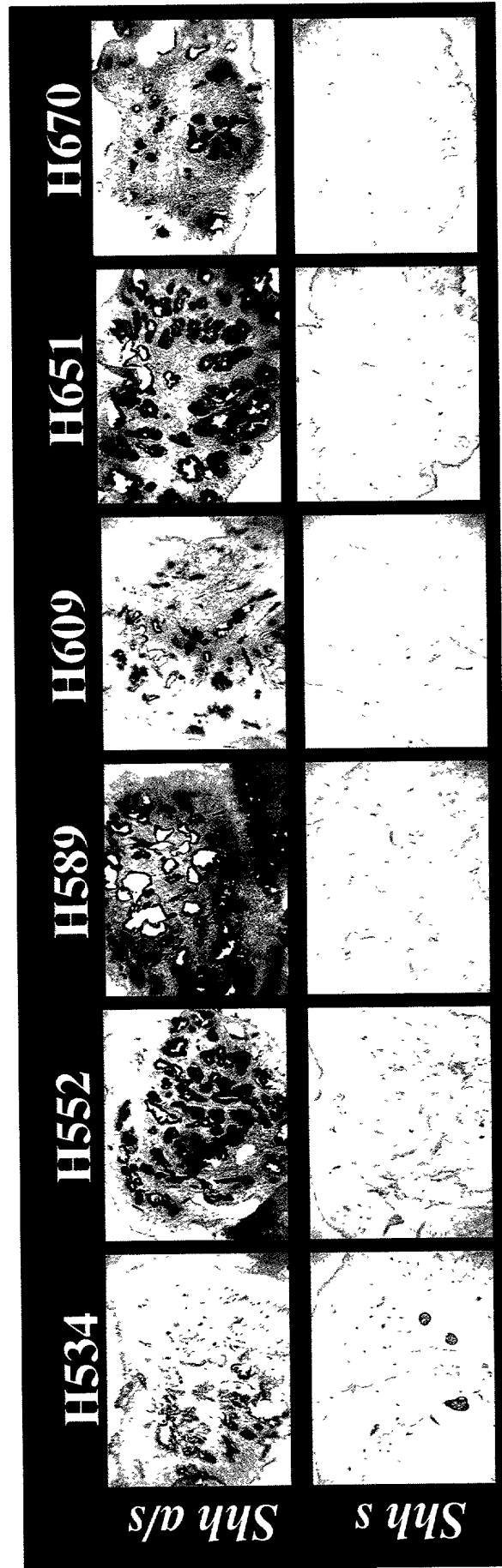


Fig. 30

Expression of *Shh* in the Prostatic Epithelium and Stroma



Patient G Patient H Patient I

Fig. 31

Expression of *Gli-1* in the Prostatic Stroma (LifeSpan)

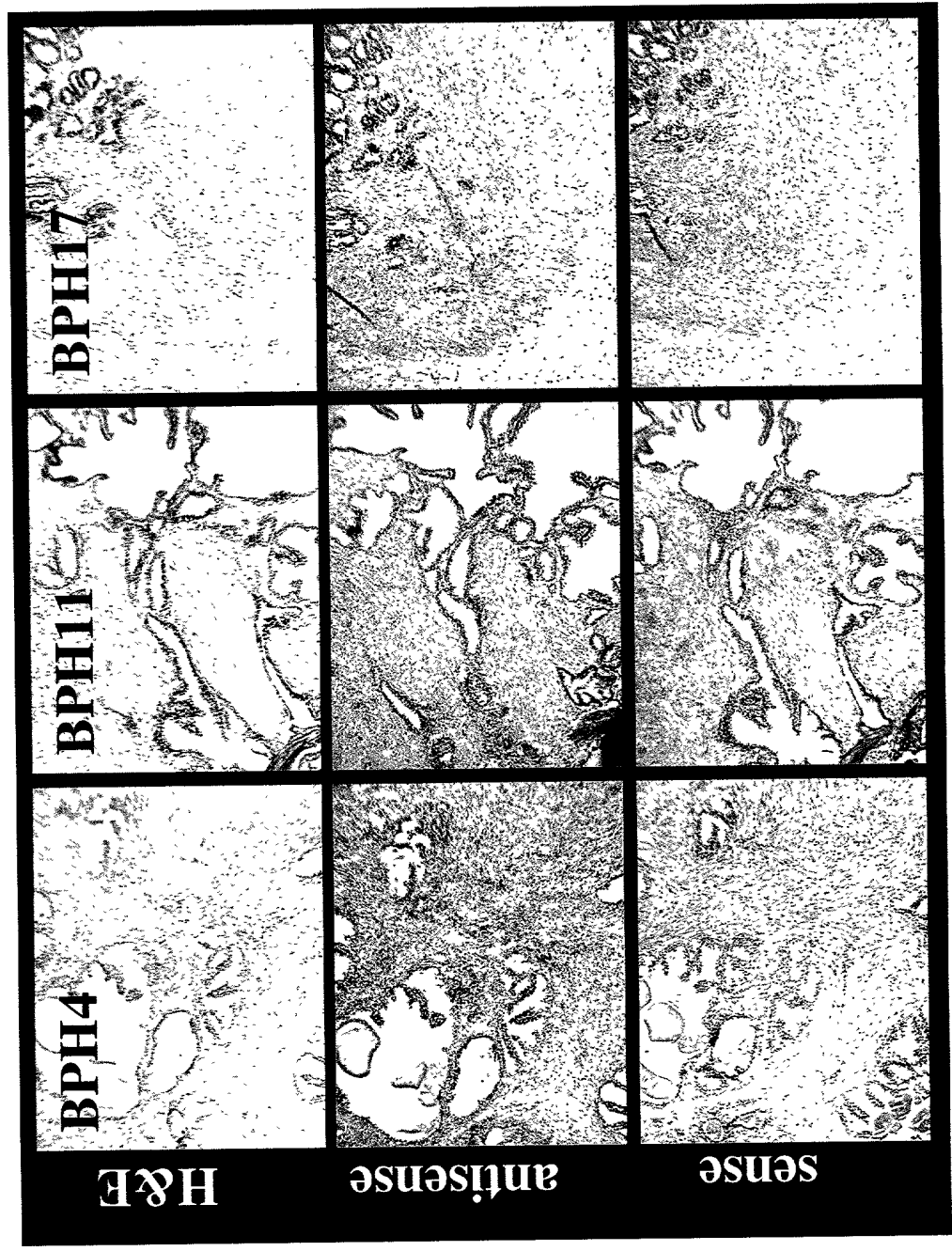


Fig. 32

Proximo-Distal Shh Gradient in Normal Prostate

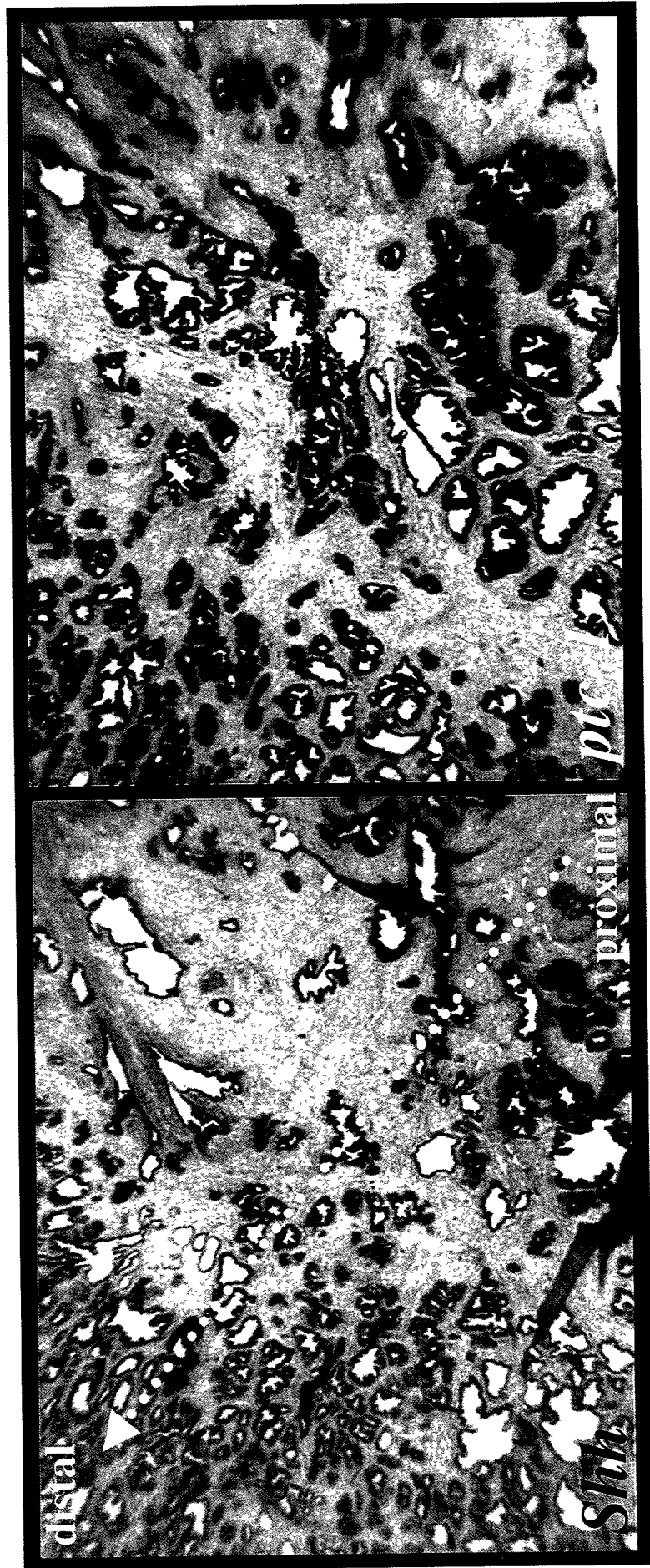


Fig. 33

Hedgehog Signaling in BPH by Q-RT-PCR

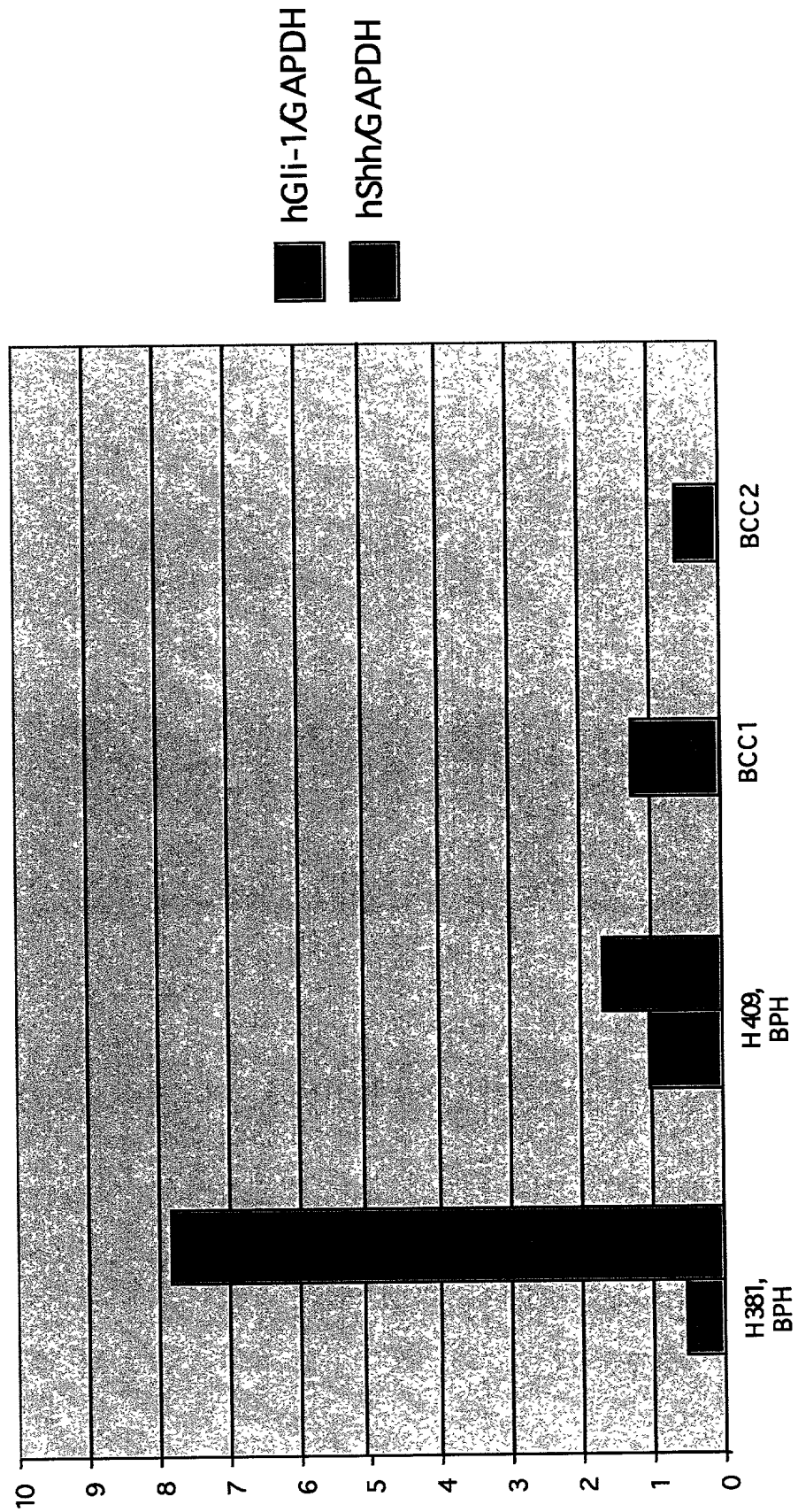
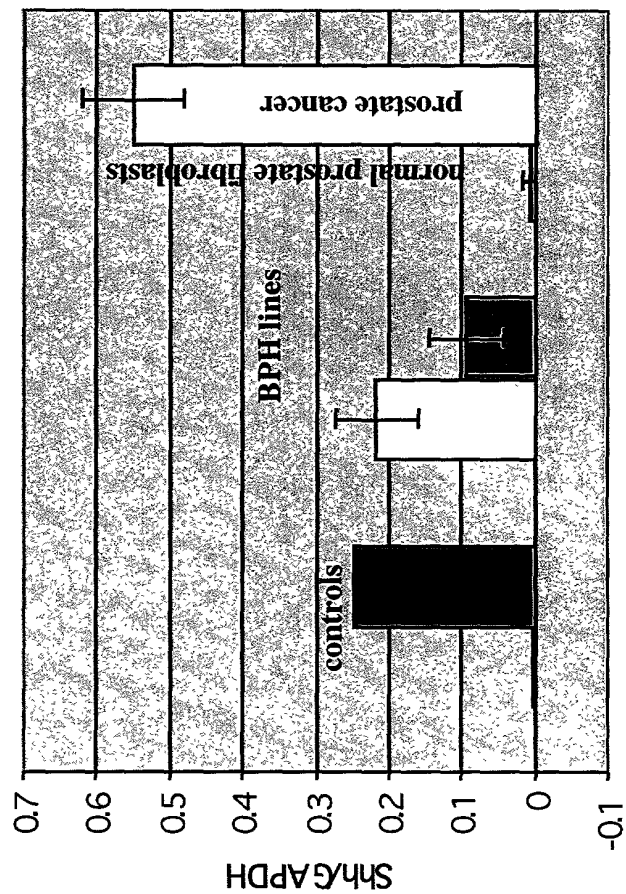


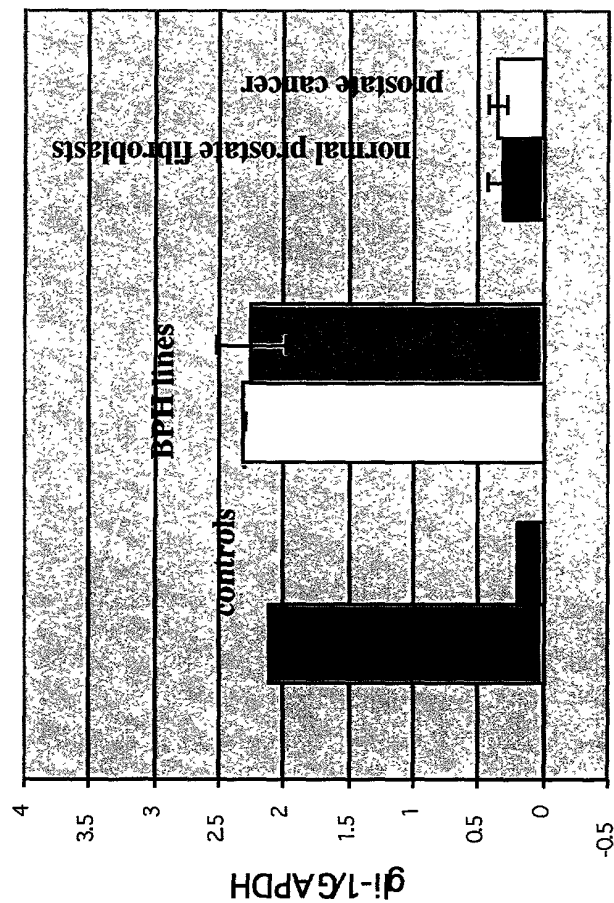
Fig. 34

HIGH LEVEL HEDGEHOG SIGNALING IN BPH CELL LINES

Shh



Gli-1



- BCC
- fetal brain
- BRFF-55T (BPH)
- 267B (BPH)
- PrSc fibroblasts
- PC-3 (prostate cancer)

Fig. 35

105T01-19822650

Effect of 5E1 on HT-29/10T1/2 Colon Cancer Growth

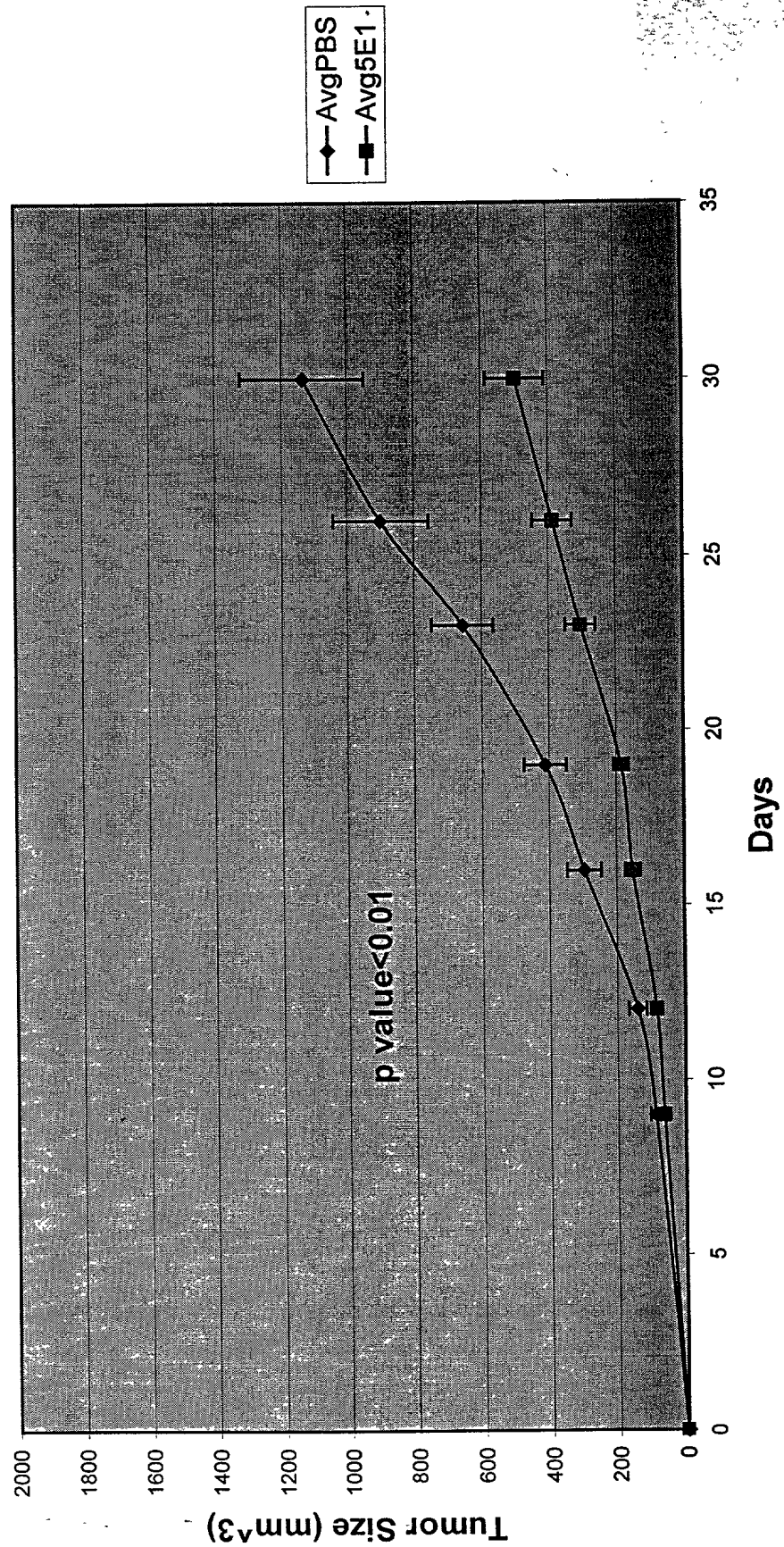


Fig. 36

Effect of 5E1 on HT-29/10T1/2 Colon Cancer Growth

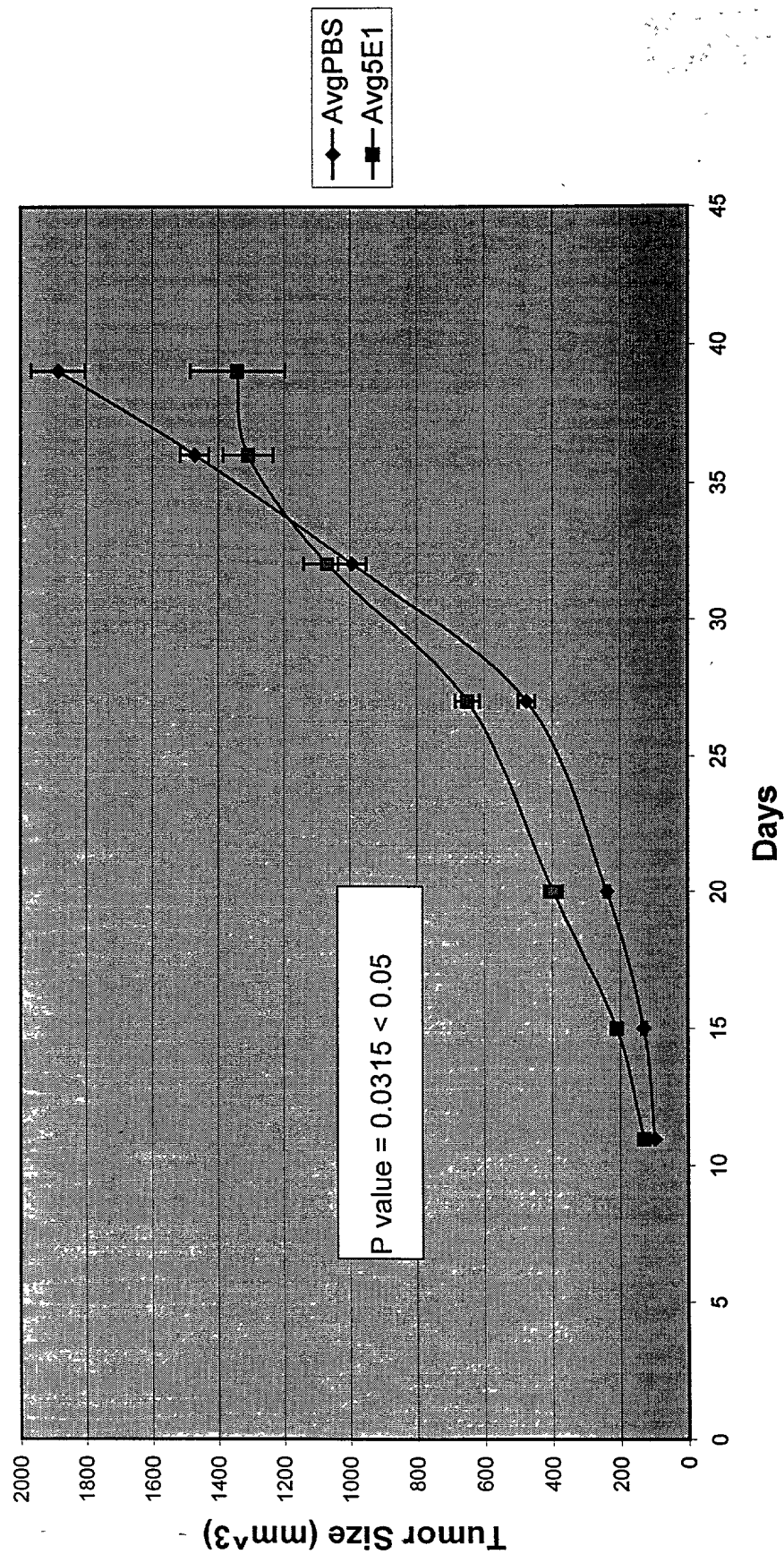


Fig. 37